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- (54) Title: HUMAN PROTEINS HAVING TRANSMEMBRANE DOMAINS AND DNAs ENCODING THESE PROTEINS
- (57) Abstract

Proteins comprising any of the amino acid sequences of SEQ ID NOS: 1 to 18 and DNAs encoding said proteins and comprising any of the nucelotide sequences of SEQ ID NOS: 19 to 36 are provided.



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DESCRIPTION

Human Proteins Having Transmembrane Domains and DNAs Encoding These Proteins

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FIELD OF THE INVENTION

The present invention relates to human proteins having transmembrane domains and cDNAs encoding these proteins. The membrane proteins of this invention can be used as pharmaceuticals or as antigens for preparing antibodies against said proteins. The cDNAs of the invention can be used as probes for the gene diagnosis and gene sources for the gene therapy. The cDNAs can also be used as gene sources for large-scale production of the membrane proteins encoded by the same. The cells into which the genes encoding the membrane proteins are introduced for expression of such membrane proteins in large amounts can be used for detection of the corresponding ligands, screening of low molecular weight medicines, etc.

20 BACKGROUND OF THE INVENTION

Membrane proteins play important roles as signal receptors, ion channels, transporters, etc. for the material transportation or information transmission mediated by the cell membrane. For instance, they are known to serve as receptors for various cytokines, ion channels for sodium ion, potassium ion, chloride ion, etc., transporters for saccharides and amino acids, and so on. The genes for many of them have been cloned already.

In recent years, it was clarified that the abnormalities

of these membrane proteins are related to a number of hitherto cryptogenic diseases. For example, a gene for a membrane protein having 12 transmembrane domains was identified as the gene responsible for cystic fibrosis [Rommens, J. M. et al., 5 Science 245: 1059-1065 (1989)]. It was also clarified that several membrane proteins act as the receptors when a virus infects the cells. For example, HIV-1 was revealed to infect into the cells through the mediation of a membrane protein fusin, a membrane protein on the T-cell membrane, having a CD-4 10 antigen and 7 transmembrane domains [Feng, Y. et al., Science 272: 872-877 (1996)]. Therefore, the discovery of new membrane proteins is anticipated to lead to the elucidation of the causes of many diseases, and the isolation of new genes coding for the membrane proteins is desired.

Heretofore, owing to the difficulty in their purification, many of membrane proteins have been isolated by an approach from the gene side. A general method is the so-called expression cloning which comprises transfection of a cDNA library in the animal cells to express the cDNA and detection 20 of the cells expressing the target membrane protein on the membrane by an immunological technique using an antibody or a physiological technique for the change in the membrane However, this method is applicable only to permeability. cloning of a gene for a membrane protein with a known function.

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In general, membrane proteins possess hydrophobic transmembrane domains inside the proteins which are synthesized in Said domains remain in the phospholipid to be the ribosome. trapped in the membrane. Accordingly, the evidence of the cDNA for encoding the membrane protein is provided by determination of the whole base sequence of a full-length cDNA and detection of highly hydrophobic transmembrane domains in the amino acid sequence of the protein encoded by said cDNA.

As a result of the extensive study, there have successful
1y been obtained human proteins having transmembrane domains,
particularly comprising any of the amino acid sequences of SEQ

ID NOS: 1 to 18, by cloning cDNAs coding for proteins having
transmembrane domains, particularly comprising any of the
nucleotide sequences of SEQ ID NOS: 19 to 36, from a human

full-length cDNA bank. The present invention is based on the
above success.

SUMMARY OF THE INVENTION

A main object of the present invention is to provide novel

human proteins having transmembrane domains, particularly
comprising any of the amino acid sequences of SEQ ID NOS: 1 to

18. Another object of this invention is to provide DNAs coding
for said novel proteins, particularly comprising any of the
nucleotide sequences of SEQ ID NOS: 19 to 36. A further object

of the invention is to provide expression vectors capable of in
vitro translating said DNAs or expressing said DNAs in
eukaryotic cells. A still further object of the invention is
to provide transformed eukaryotic cells capable of expressing
said DNAs to produce said proteins.

In one embodiment, the present invention provides a composition comprising a protein, wherein said protein comprises an amino acid sequence selected from the group consisting of the amino acid sequences of SEQ ID NOS: 1 to 18 and their fragments.

In another embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 19 to 36.

In a further embodiment, the present invention provides a composition comprising an isolated polynucleotide selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 37 to 54.

10 BRIEF DESCRIPTION OF DRAWINGS

- Figure 1: A figure depicting the structure of the secretory signal sequence detection vector pSSD3.
- Figure 2: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01263.
- 15 Figure 3: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01299.
 - Figure 4: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01347.
- Figure 5: A figure depicting the hydrophobicity/hydrophi-20 licity profile of the protein encoded by clone HP01440.
 - Figure 6: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP01526.
 - Figure 7: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10230.
- 25 Figure 8: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10389.
 - Figure 9: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10408.
 - Figure 10: A figure depicting the hydrophobicity/hydro-

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philicity profile of the protein encoded by clone HP10412.

Figure 11: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10413.

Figure 12: A figure depicting the hydrophobicity/hydro-5 philicity profile of the protein encoded by clone HP10415.

Figure 13: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10419.

Figure 14: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10424.

Figure 15: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10428.

Figure 16: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10429.

Figure 17: A figure depicting the hydrophobicity/hydro-15 philicity profile of the protein encoded by clone HP10432.

Figure 18: A figure depicting the hydrophobicity/hydro-philicity profile of the protein encoded by clone HP10433.

Figure 19: A figure depicting the hydrophobicity/hydrophilicity profile of the protein encoded by clone HP10480.

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BEST MODE FOR CARRING OUT INVENTION

The proteins of the present invention can be obtained, for example, by isolation from human organs, cell lines, etc., by chemical synthesis on the basis of the amino acid sequences as herein disclosed, or by recombinant DNA technology using the DNA encoding the transmembrane domains of the invention. Among them, adoption of the recombinant DNA technology is preferred. Specifically, each of the proteins may be prepared by in vitro transcription of a vector comprising the cDNA of the invention

to make RNA and in vitro translation using this RNA as a template to accomplish in vitro expression. Also, each of the proteins may be prepared in a large amount by the use of Escherichia coli, Bacillus subtilis, yeasts, animal cells, etc. comprising a suitable expression vector having the DNA encoding such protein.

In the case of producing the protein of the invention by the use of a microorganism such as Escherichia coli, the translation region of the cDNA of the invention is constructed 10 in an expression vector having an origin, a promoter, a ribosome-binding site, a cDNA-cloning site, a terminator, etc. that can be replicated in the microorganism and, after transformation of the host cells with said expression vector, the resultant transformant is incubated, whereby the protein 15 encoded by said cDNA can be produced in a large amount in the microorganism. In that case, a protein fragment containing an optional region can be obtained by performing the expression with inserting an initiation codon and a termination codon before and after the optional translation region. Alternative-20 ly, a fusion protein with another protein can be expressed. Only a protein portion encoding said cDNA can be obtained by cleavage of said fusion protein with an appropriate protease.

For production of the protein of the invention by expression of DNA coding for such protein in eukaryotic cells, the translation region of said cDNA may be recombined into an expression vector for eukaryotic cells having a promoter, a splicing domain, a poly(A) addition site, etc., followed by introduction into eukaryotic cells so that the protein of the invention is produced as a membrane protein on the cell

membrane surface. Examples of the expression vector are pKA1, pED6_dpc2, pCDM8, pSVK3, pMSG, pSVL, pBK-CMV, pBK-RSV, EBV vector, pRS, pYES2, etc. As the eukaryotic cells, there are exemplified mammalian animal culture cells (e.g. simian kidney cells COS7, chinese hamster ovary cells CHO), budding yeasts, Schizosaccharomyces pombe, silkworm cells, Xenopus laevis egg cells, etc., but any other eukaryotic cells may also be used insofar as the protein of the invention can be expressed on the membrane surface. In order to introduce the expression vector into eukaryotic cells, there may be adopted any conventional procedure such as electroporation, calcium phosphate method, liposome method or DEAE dextran method.

The proteins of the present invention include peptide fragments (5 or more amino acid residues) containing any 15 partial amino acid sequence of the amino acid sequences of SEQ ID NOS: 1 to 18. These fragments can be used as antigens for Also, the proteins of the preparation of the antibodies. invention that have signal sequences appear in the form of maturation proteins on the cell surface, after the signal 20 sequences are removed. Therefore, these maturation proteins shall come within the scope of the present invention. The Nterminal amino acid sequences of the maturation proteins can be easily identified by using the method for the cleavage-site determination in a signal sequence [Japan Patent Kokai No. 187100/96]. Further, many membrane proteins are subjected to the processing on the cell surface to be converted to the secretor forms. These secretor proteins or peptides shall come within the scope of the present invention. When glycosylation sites are present in the amino acid sequences, expression in appropriate animal cells affords glycosylated proteins. Therefore, these glycosylated proteins or peptides also shall come within the scope of the invention.

The DNAs of the invention include all DNAs encoding the 5 above-mentioned proteins. Said DNAs can be obtained using the method by chemical synthesis, the method by cDNA cloning, and so on.

Each of the cDNAs of the invention can be cloned from, for example, the cDNA libraries of the human cell origin. The cDNA 10 is synthesized using as a template a poly(A) + RNA extracted from human cells. The human cells may be cells delivered from the human body, for example, by the operation or may be the culture cells. The cDNA can be synthesized by using any method selected from the Okayama-Berg method [Okayama, H. and Berg, 15 P., Mol. Cell. Biol. 2: 161-170 (1982)], the Gubler-Hoffman method [Gubler, U. and Hoffman, J. Gene 25: 263-269 (1983)], and so on, but it is preferred to use the capping method [Kato, S. et al., Gene 150: 243-250 (1994)] as illustrated in Examples in order to obtain a full-length clone in an effective manner.

The primary selection of a cDNA encoding a human protein having transmembrane domains is performed by the sequencing of a partial base sequence of the cDNA clone selected at random from the cDNA libraries, sequencing of the amino acid sequence encoded by the base sequence, and recognition of the presence 25 or absence of hydrophobic site(s) in the resulting N-terminal amino acid sequence region. Next, the secondary selection is carried out by determination of the whole base sequence by the sequencing and the protein expression by the in vitro translation. The ascertainment of the cDNA of the present

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invention for encoding the protein having the secretory signal sequence is performed by using the signal sequence detection method [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)]. In other words, the ascertainment for the coding portion of the inserted cDNA fragment to function as a signal sequence is provided by fusing a cDNA fragment encoding the N-terminus of the target protein with a cDNA encoding the protease domain of urokinase and then expressing the resulting cDNA in COS7 cells to detect the urokinase activity in the cell culture medium. On the other hand, the N-terminal region is judged to remain in the membrane in the case where the urokinase activity is not detected in the cell culture medium.

The cDNAs of the invention are characterized by containing any of the nucleotide sequences of SEQ ID NOS: 19 to 36 or any of the nucleotide sequences of SEQ ID NOS: 37 to 54. Table 1 summarizes the clone number (HP number), the cells affording the cDNA, the total nucleotide number of the cDNA, and the number of the amino acid residues of the encoded protein, for each of the cDNAs.

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Table 1

5	Sequence HP		Cells	Number of Nucleotides	Number of Amino Acid
					Residues
10	1, 19, 37	HP01263	Liver	1502	382
	2, 20, 38	HP01299	Liver	1349	317
	3, 21, 39	HP01347	Liver	1643	296
	4, 22, 40	HP01440	Stomach cancer	729	197
	5, 23, 41	HP01526	Stomach cancer	1322	221
20	6, 24, 42	HP10230	Stomach cancer	. 3045	251
	7, 25, 43	HP10389	КВ	653	106
	8, 26, 44	HP10408	Stomach cancer	439	78
25	9, 27, 45	HP10412	Stomach cancer	1131	314
	10, 28, 46	HP10413	Stomach cancer	1875	195
30	11, 29, 47	HP10415	Stomach cancer	1563	462
	12, 30, 48	HP10419	Stomach cancer	2030	247
	13, 31, 49	HP10424	Stomach cancer	493	113
35	14, 32, 50	HP10428	КВ	2044	365
	15, 33, 51	HP10429 ·	Stomach cancer	1043	226
40	16, 34, 52	HP10432	Liver	972	129
	17, 35, 53	HP10433	Liver	695	163
	18, 36, 54	HP10480	Stomach cancer	1914	193

Hereupon, the same clone as any of the cDNAs of the invention can be easily obtained by screening of the cDNA libraries constructed from the cell line or the human tissues employed in the invention, by the use of an oligonucleotide probe synthesized on the basis of the corresponding cDNA nucleotide sequence of SEQ ID NOS: 37 to 54.

In general, the polymorphism due to the individual difference is frequently observed in human genes. Therefore, any cDNA that is subjected to insertion or deletion of one or plural nucleotides and/or substitution with other nucleotides

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in SEQ ID NOS: 37 to 54 shall come within the scope of the invention.

In a similar manner, any protein that is produced by these modifications comprising insertion or deletion of one or plural nucleotides and/or substitution with other nucleotides shall come within the scope of the present invention, as far as said protein possesses the activity of the corresponding protein having the amino acid sequence of SEQ ID NOS: 1 to 18.

The cDNAs of the invention include cDNA fragments (more than 10 bp) containing any partial nucleotide sequence of the nucleotide sequence of SEQ ID NOS: 19 to 36 or of the nucleotide sequence of SEQ ID NOS: 37 to 54. Also, DNA fragments consisting of a sense chain and an anti-sense chain shall come within this scope. These DNA fragments can be used as the probes for the gene diagnosis.

The present invention also provides genes corresponding to the polynucleotide sequences disclosed herein. "Corresponding genes" are the regions of the genome that are transcribed to produce the mRNAs from which cDNA polynucleotide sequences are derived and may include contiguous regions of the genome necessary for the regulated expression of such genes. Corresponding genes may therefore include but are not limited to coding sequences, 5' and 3' untranslated regions, alternatively spliced exons, introns, promoters, enhancers, and silencer or suppressor elements. The corresponding genes can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include the preparation of probes or primers from the disclosed sequence information for identification and/or amplification of genes in appropriate

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genomic libraries or other sources of genomic materials. "isolated gene" is a gene that has been separated from the adjacent coding sequences, if any, present in the genome of the organism from which the gene was isolated.

Organisms that have enhanced, reduced, or modified expression of the gene(s) corresponding to the polynucleotide sequences disclosed herein are provided. The desired change in gene expression can be achieved through the use of antisense polynucleotides or ribozymes that bind and/or cleave 10 the mRNA transcribed from the gene (Albert and Morris, 1994, Trends Pharmacol. Sci. 15(7): 250-254; Lavarosky et al., 1997, Biochem. Mol. Med. 62(1): 11-22; and Hampel, 1998, Prog. Nucleic Acid Res. Mol. Biol. 58: 1-39; all of which are incorporated by reference herein). Transgenic animals that 15 have multiple copies of the gene(s) corresponding to the polynucleotide sequences disclosed herein, preferably produced by transformation of cells with genetic constructs that are stably maintained within the transformed cells and their Transgenic animals that have modified progeny, are provided. 20 genetic control regions that increase or reduce gene expression levels, or that change temporal or spatial patterns of gene expression, are also provided (see European Patent No. 0 649 In addition, 464 B1, incorporated by reference herein). organisms are provided in which the gene(s) corresponding to 25 the polynucleotide sequences disclosed herein have been partially or completely inactivated, through insertion of extraneous sequences into the corresponding gene(s) or through deletion of all or part of the corresponding gene(s). Partial or complete gene inactivation can be accomplished through

insertion, preferably followed by imprecise excision, of transposable elements (Plasterk, 1992, Bioessays 14(9): 629-633; Zwaal et al., 1993, Proc. Natl. Acad. Sci. USA 90(16): 7431-7435; Clark et al., 1994, Proc. Natl. Acad. Sci. USA 5 91(2): 719-722; all of which are incorporated by reference herein), or through homologous recombination, preferably detected by positive/negative genetic selection strategies (Mansour et al., 1988, Nature 336: 348-352; U.S. Patent Nos. 5,464,764; 5,487,992; 5,627,059; 5,631,153; 5,614, 10 5,616,491; and 5,679,523; all of which are incorporated by These organisms with altered gene reference herein). expression are preferably eukaryotes and more preferably are mammals. Such organisms are useful for the development of non-human models for the study of disorders involving the 15 corresponding gene(s), and for the development of assay systems for the identi fication of molecules that interact with the protein product(s) of the corresponding gene(s).

Where the protein of the present invention is membrane-bound (e.g., is a receptor), the present invention also provides for soluble forms of such protein. In such forms part or all of the intracellular and transmembrane domains of the protein are deleted such that the protein is fully secreted from the cell in which it is expressed. The intracellular and transmembrane domains of proteins of the invention can be identified in accordance with known techniques for determination of such domains from sequence information.

Proteins and protein fragments of the present invention include proteins with amino acid sequence lengths that are at least 25% (more preferably at least 50%, and most preferably at

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least 75%) of the length of a disclosed protein and have at least 60% sequence identity (more preferably, at least 75% identity; most preferably at least 90% or 95% identity) with that disclosed protein, where sequence identity is determined 5 by comparing the amino acid sequences of the proteins when aligned so as to maximize overlap and identity while minimizing sequence gaps. Also included in the present invention are proteins and protein fragments that contain a segment preferably comprising 8 or more (more preferably 20 or more, 10 most preferably 30 or more) contiguous amino acids that shares at least 75% sequence identity (more preferably, at least 85% identity; most preferably at least 95% identity) with any such segment of any of the disclosed proteins.

Species homologs of the disclosed polynucleotides and proteins are also provided by the present invention. As used herein, a "species homologue" is a protein or polynucleotide with a different species of origin from that of a given protein or polynucleotide, but with significant sequence similarity to the given protein or polynucleotide, as determined by those of skill in the art. Species homologs may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source from the desired species.

The invention also encompasses allelic variants of the 25 disclosed polynucleotides or proteins; that is, naturally-occurring alternative forms of the isolated polynucleotide which also encode proteins which are identical, homologous, or related to that encoded by the polynucleotides.

The invention also includes polynucleotides with sequences

complementary to those of the polynucleotides disclosed herein.

The present invention also includes polynucleotides capable of hybridizing under reduced stringency conditions, more preferably stringent conditions, and most preferably 5 highly stringent conditions, to polynucleotides described herein. Examples of stringency conditions are shown in the table below: highly stringent conditions are those that are at least as stringent as, for example, conditions A-F; stringent conditions are at least as stringent as, for example, conditions G-L; and reduced stringency conditions are at least as stringent as, for example,

Table 2

Stringency	Polynucleotide	Hybrid	Hybridization Temperature	Wash
Condition	Hybrid	Length	and Buffer [†]	Temperature
• • • • • • • • • • • • • • • • • • • •	,	(bp) [‡]		and Buffer [†]
A	DNA : DNA	≥50	65°C; 1×SSC -or-	65°C; 0.3×SSC
			42°C; 1×SSC,50% formamide	
В	DNA : DNA	<50	T _B *; 1×SSC	T_B^* ; 1×SSC
C	DNA : RNA	≥50	67℃; 1×SSC -or-	67°C; 0.3×SSC
			45°C; 1×SSC,50% formamide	
D	DNA : RNA	<50	T _D *; 1×SSC	T _D *; 1×SSC
E	RNA : RNA	≥50	70°C; 1×SSC -or-	70℃; 0.3×SSC
			50°C; 1×SSC,50% formamide	
F	RNA : RNA	<50	T _F *; 1×SSC	T _F *; 1×SSC
G	DNA : DNA	≥50	65°C; 4×SSC -or-	65℃; 1×SSC
			42°C; 4×SSC,50% formamide	
H	DNA : DNA	<50	T _H *; 4×SSC	T _H *; 4×SSC
I	DNA : RNA	≥50.	67°C; 4×SSC -or-	67℃; 1×SSC
			45℃; 4×SSC,50% formamide	
J	DNA : RNA	<50	T_J^* ; 4×SSC	T _J *; 4×SSC
K	RNA: RNA	≥50	70°C; 4×SSC -or-	67℃; 1×SSC
			50℃; 4×SSC,50% formamide	
L	RNA: RNA	<50	T _L *; 2×SSC	T_L^* ; 2×SSC
M	DNA : DNA	≥50	50℃; 4×SSC -or-	50°C; 2×SSC
			40℃; 6×SSC,50% formamide	
N	DNA : DNA	<50	T _N *; 6×SSC	T _N *; 6×SSC
0	DNA : RNA	≥50	55°C; 4×SSC -or-	55℃; 2×SSC
			42℃; 6×SSC,50% formamide	
P	DNA : RNA	<50	T _P *; 6×SSC	T _P *; 6×SSC
Q	RNA : RNA	≥50	60℃; 4×SSC -or-	60°C; 2×SSC
			45℃; 6×SSC,50% formamide	
R	RNA : RNA	<50	T_R^* ; 4×SSC	T _R *; 4×SSC

- ‡: The hybrid length is that anticipated for the hybridized region(s) of the hybridizing polynucleotides. When hybridizing a polynucleotide to a target polynucleotide of unknown sequence, the hybrid length is assumed to be that of the hybridizing polynucleotide. When polynucleotides of known sequence are hybridized, the hybrid length can be determined by aligning the sequences of the polynucleotides and identifying the region or regions of optimal sequence complementarity.
- \dagger : SSPE (1×SSPE is 0.15M NaCl, 10mM NaH₂PO₄, and 1.25mM EDTA, pH7.4) can be substituted for SSC (1×SSC is 0.15M NaCl and 15mM sodium citrate) in the hybridization and wash buffers; washes are performed for 15 minutes after hybridization is complete.
- * T_B T_R : The hybridization temperature for hybrids anticipated to be less than 50 base pairs in length should be 5-10°C less than the melting temperature (T_m) of the hybrid, where T_m is determined according to the following equations. For hybrids less than 18 base pairs in length, T_m (°C)=2(#of A + T bases) + 4(# of G + C bases). For hybrids between 18 and 49 base pairs in length, T_m (°C)=81.5 + 16.6(log₁₀[Na⁺]) + 0.41 (%G+C) (600/N), where N is the number of bases in the hybrid, and [Na⁺] is the concentration of sodium ions in the hybridization buffer ([Na⁺] for 1×SSC=0.165M).

Additional examples of stringency conditions for polynucleotide hybridization are provided in Sambrook, J., E.F. Fritsch, and T. Maniatis, 1989, Molecular Cloning: A Laboratory

- 5 Manual, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, chapters 9 and 11, and Current Protocols in Molecular Biology, 1995, F.M. Ausubel et al., eds., John Wiley & Sons, Inc., sections 2.10 and 6.3-6.4, incorporated herein by reference.
- length that is at least 25% (more

 preferably at least 50%, and most preferably at least 75%) of
 the length of the polynucleotide of
 the present invention to which it hybridizes, and has at least

 60% sequence identity (more
 preferably, at least 75% identity; most preferably at least 90%
 or 95% identity) with the
 polynucleotide of the present invention to which it hybridizes,
 where sequence identity is
- 20 determined by comparing the sequences of the hybridizing polynucleotides when aligned so as to maximize overlap and identity while minimizing sequence gaps.

25 EXAMPLE

The present invention is embodied in more detail by the following examples, but this embodiment is not intended to restrict the present invention. The basic operations and the enzyme reactions with regard to the DNA recombination are

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carried out according to the literature ["Molecular Cloning. A Laboratory Manual", Cold Spring Harbor Laboratory, 1989]. Unless otherwise stated, restrictive enzymes and a variety of modification enzymes to be used were those available from 5 Takara Shuzo Co., Ltd. The manufacturer's instructions were used for the buffer compositions as well as for the reaction conditions, in each of the enzyme reactions. The cDNA synthesis was carried out according to the literature [Kato, S. et al., Gene 150: 243-250 (1994)].

(1) Preparation of Poly(A) + RNA

The epidermoid carcinoma cell line KB (ATCC CRL 17), tissues of stomach cancer delivered by the operation, and liver were used for human cells to extract mRNAs. The cell line was cultured by a conventional procedure.

After about 1 g of human tissues was homogenized in 20 ml of a 5.5 M guanidinium thiocyanate solution, total mRNAs were prepared in accordance with the literature [Okayama, H. et al., "Methods in Enzymology" Vol. 164, Academic Press, 1987]. These mRNAs were subjected to chromatography using an oligo(dT)-20 cellulose column washed with 20 mM Tris-hydrochloric acid buffer solution (pH 7.6), 0.5 M NaCl, and 1 mM EDTA to obtain a poly(A) + RNA in accordance with the above-mentioned literature.

(2) Construction of cDNA Library

To a solution of 10 μg of the above-mentioned poly(A)⁺ RNA 25 in 100 mM Tris-hydrochloric acid buffer solution (pH 8) was added one unit of an RNase-free, bacterium-origin alkaline phosphatase and the resulting solution was allowed to react at 37°C for one hour. After the reaction solution underwent the

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phenol extraction followed by the ethanol precipitation, the obtained pellets were dissolved in a mixed solution of 50 mM sodium acetate (pH 6), 1 mM EDTA, 0.1% 2-mercaptoethanol, and 0.01% Triton X-100. Thereto was added one unit of a tobacco-5 origin pyrophosphatase (Epicenter Technologies) and the resulting solution at a total volume of 100 μl was allowed to react at 37°C for one hour. After the reaction solution underwent the phenol extraction followed by the ethanol precipitation, the thus-obtained pellets were dissolved in 10 water to obtain a decapped poly(A) + RNA solution.

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To a solution of the decapped $poly(A)^+$ RNA and 3 nmol of a DNA-RNA chimeric oligonucleotide (5'-dG-dG-dG-dG-dA-dA-dT-dTdC-dG-dA-G-G-A-3') in a mixed aqueous solution of 50 mM Trishydrochloric acid buffer solution (pH 7.5), 0.5 mM ATP, 5 mM $\,$ 15 $MgCl_2$, 10 mM 2-mercaptoethanol, and 25% polyethylene glycol were added 50 units of T4 RNA ligase and the resulting solution at a total volume of 30 μl was allowed to react at 20°C for 12 hours. After the reaction solution underwent the phenol extraction followed by the ethanol precipitation, the thus-20 obtained pellets were dissolved in water to obtain a chimeric oligo-capped poly(A) + RNA.

After the vector pKAl developed by the present inventors (Japanese Patent Kokai Publication No. 1992-117292) was digested with KpnI, an about 60-dT tail was inserted by a 25 terminal transferase. This product was digested with EcoRV to remove the dT tail at one side and the resulting molecule was used as a vectorial primer.

After 6 µg of the previously-prepared chimeric oligocapped poly(A) $^+$ RNA was annealed with 1.2 μ g of the vectorial

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primer, the product was dissolved in a mixed solution of 50 mM Tris-hydrochloric acid buffer solution (pH 8.3), 75 mM KCl, 3 mM ${\rm MgCl}_2$, 10 mM dithiothreitol, and 1.25 mM dNTP (dATP + dCTP + dGTP + dTTP), mixed with 200 units of a reverse transferase 5 (GIBCO-BRL), and the resulting solution at a total volume of 20 μl was allowed to react at 42°C for one hour. After the reaction solution underwent the phenol extraction followed by the ethanol precipitation, the thus-obtained pellets were dissolved in a mixed solution of 50 mM Tris-hydrochloric acid 10 buffer solution (pH 7.5), 100 mM NaCl, 10 mM MgCl2, and 1 mM dithiothreitol. Thereto were added 100 units of EcoRI and the resulting solution at a total volume of 20 μl was allowed to react at 37°C for one hour. After the reaction solution underwent the phenol extraction followed by the ethanol 15 precipitation, the obtained pellets were dissolved in a mixed solution of 20 mM Tris-hydrochloric acid buffer solution (pH 7.5), 100 mM KCl, 4 mM MgCl₂, 10 mM $(NH_4)_2SO_4$, and 50 $\mu g/ml$ Thereto were added 60 units of bovine serum albumin. Escherichia coli DNA ligase and the resulting solution was 20 allowed to react at 16°C for 16 hours. To the reaction solution were added 2 µl of 2 mM dNTP, 4 units of Escherichia coli DNA polymerase I, and 0.1 unit of Escherichia coli DNase H and the resulting solution was allowed to react at 12°C for one hour and then at 22°C for one hour.

Next, the cDNA-synthesis reaction solution was used to transform <code>Escherichia coli DH12S</code> (GIBCO-BRL). The transformation was carried out by the electroporation method. A portion of the transformant was inoculated on a 2xYT agar culture medium containing 100 μ g/ml ampicillin, which was

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incubated at 37°C overnight. A colony grown on the culture medium was randomly picked up and inoculated on 2 ml of the 2xYT culture medium containing 100 $\mu\text{g/ml}$ ampicillin, which was incubated at 37°C overnight. The culture medium was centrifuged 5 to separate the cells, from which a plasmid DNA was prepared by the alkaline lysis method. After the plasmid DNA was doubledigested with EcoRI and NotI, the product was subjected to 0.8% agarose gel electrophoresis to determine the size of the cDNA insert. In addition, by the use of the obtained plasmid as a 10 template, the sequence reaction using M13 universal primer labeled with a fluorescent dye and Taq polymerase (a kit of Applied Biosystems Inc.) was carried out and the product was analyzed by a fluorescent DNA-sequencer (Applied Biosystems Inc.) to determine the base sequence of the cDNA 5'-terminal of about 400 bp. The sequence data were filed as a homo-protein cDNA bank data base.

(3) Selection of cDNAs Encoding Proteins Having
Transmembrane Domains

The base sequence registered in the homo-protein cDNA bank

20 data base was converted to three frames of amino acid sequences
and the presence or absence of an open reading frame (ORF)
beginning from the initiation codon. Then, the selection was
made for the presence of a signal sequence that is
characteristic to a secretory protein at the N-terminal of the

25 portion encoded by ORF. These clones were sequenced from the
both 5' and 3' directions by using the deletion method to
determine the sequence of the whole base sequence. The
hydrophobicity/hydrophilicity profiles were obtained for
proteins encoded by ORF by the Kyte-Doolittle method [Kyte, J.

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& Doolittle, R. F., J. Mol. Bio. 157: 105-132 (1982)] to examine the presence or absence of a hydrophobic region. In the case in which there is a hydrophobic region of putative transmembrane domain(s) in the amino acid sequence of an encoded protein, this protein was considered as a membrane protein.

(4) Construction of Secretory Signal Detection Vector pSSD3

One microgram of pSSD1 carrying the SV40 promoter and a cDNA encoding the protease domain of urokinase [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)] was digested with 5 units of Bg1II and 5 units of EcoRV. Then, after dephosphorylation at the 5' terminal by the CIP treatment, a DNA fragment of about 4.2 kbp was purified by cutting off from the gel of agarose gel electrophoresis.

Two oligo DNA linkers, L1 (5'-GATCCCGGGTCACGTGGGAT-3') and synthesized L2 (5'-ATCCCACGTGACCCGG-3'), were phosphorylated by T4 polynucleotide kinase. After annealing of the both linkers, followed by ligation with the previously-20 prepared pSSD1 fragment by T4 DNA ligase, Escherichia coli JM109 was transformed. A plasmid pSSD3 was prepared from the transformant and the objective recombinant was confirmed by the determination of the base sequence of the linker-inserted fragment. Figure 1 illustrates the structure of the thus-25 obtained plasmid. The present plasmid vector carries three types of blunt-end formation restriction enzyme sites, SmaI, PmaCI, and EcoRV. Since these cleavage sites are positioned in succession at an interval of 7 bp, selection of an appropriate site in combination of three types of frames for the inserting

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cDNA allows to construct a vector expressing a fusion protein.

(5) Functional Verification of Secretory Signal Sequence Whether the N-terminal hydrophobic region in the secretory protein clone candidate obtained in the above-mentioned steps 5 functions as the secretory signal sequence was verified by the method described in the literature [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)]. First, the plasmid containing the target cDNA was cleaved at an appropriate restriction enzyme site that existed at the downstream of the portion 10 expected for encoding the secretory signal sequence. In the case in which this restriction enzyme site was a protruding terminus, the site was blunt-ended by the Klenow treatment or treatment with the mung-bean nuclease. Digestion with HindIII was further carried out and a DNA fragment containing the SV40 15 promoter and a cDNA encoding the secretory sequence at the downstream of the promoter was separated by agarose gel electrophoresis. This fragment was inserted between the pSSD3 HindIII site and a restriction enzyme site selected so as to match with the urokinase-coding frame, thereby constructing a 20 vector expressing a fusion protein of the secretory signal portion of the target cDNA and the urokinase protease domain.

After Escherichia coli (host: JM109) bearing the fusionprotein expression vector was incubated at 37°C for 2 hours in 2 ml of the 2xYT culture medium containing 100 $\mu g/ml$ 25 ampicillin, the helper phage M13KO7 (50 μ l) was added and the incubation was continued at 37°C overnight. A supernatant separated by centrifugation underwent precipitation with polyethylene glycol to obtain single-stranded phage particles. These particles were suspended in 100 µl of 1 mM Tris-0.1 mM

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EDTA, pH 8 (TE). Also, there was used as a control a suspension of single-stranded particles prepared in the same manner from the vector pLA1-UPA containing pSSD3 and a full-length cDNA of urokinase [Yokoyama-Kobayashi, M. et al., Gene 163: 193-196 (1995)].

simian-kidney-origin culture cells, COS7, incubated at $37\,^{\circ}\mathrm{C}$ in the presence of 5% CO_2 in the Dulbecco's modified Eagle's culture medium (DMEM) containing 10% bovine fetus albumin. Into a 6-well plate (Nunc Inc., 3 cm in the well 10 diameter) were inoculated 1 \times 10 5 COS7 cells and incubation was carried out at 37°C for 22 hours in the presence of 5% $\rm CO_2$. After the culture medium was removed, the cell surface was washed with a phosphate buffer solution and then washed again with DMEM containing 50 mM Tris-hydrochloric acid (pH 7.5) 15 (TDMEM). To the cells were added 1 μ l of the single-stranded phage suspension, 0.6 ml of the DMEM culture medium, and 3 μ l of TRANSFECTAMTM (IBF Inc.) and the resulting mixture was incubated at 37°C for 3 hours in the presence of 5% CO2. After the sample solution was removed, the cell surface was washed 20 with TDMEM, 2 ml per well of DMEM containing 10% bovine fetus albumin was added, and the incubation was carried out at 37°C for 2 days in the presence of 5% CO2.

To 10 ml of 50 mM phosphate buffer solution (pH 7.4) containing 2% bovine fibrinogen (Miles Inc.), 0.5% agarose, and 1 mM potassium chloride were added 10 units of human thrombin (Mochida Pharmaceutical Co., Ltd.) and the resulting mixture was solidified in a plate of 9 cm in diameter to prepare a fibrin plate. Ten microliters of the culture supernatant of the

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transfected COS7 cells were spotted on the fibrin plate, which was incubated at 37°C for 15 hours. The diameter of the thusobtained clear circle was taken as an index for the urokinase activity. In the case in which a cDNA fragment codes for the 5 amino acid sequence that functions as a secretory signal sequence, a fusion protein is secreted to form a clear circle by its urokinase activity. Therefore, in the case in which a clear circle is not formed, the fusion protein remains as trapped in the membrane and the cDNA fragment is considered to 10 code for a transmembrane domain.

(6) Protein Synthesis by In Vitro Translation

The plasmid vector carrying the cDNA of the present invention was utilized for the transcription/translation by the $T_{N}T$ rabbit reticulocyte lysate kit (Promega Biotec). In this 15 case, [35]methionine was added and the expression product was labeled with the radioisotope. All reactions were carried out by following the protocols attached to the kit. Two micrograms of the plasmid was allowed to react at 30°C for 90 minutes in total 25 ml of a reaction solution containing 12.5 μl of the 20 $\ensuremath{T_{N}T}$ rabbit reticulocyte lysate, 0.5 μl of the buffer solution (attached to the kit), 2 μl of an amino acid mixture (methionine-free), 2 μl (0.37 MBq/ μl) of [^{35}S]methionine (Amersham Corporation), 0.5 μl of T7 RNA polymerase, and 20 U of RNasin. To 3 μl of the reaction solution was added 2 μl of 25 an SDS sampling buffer (125 mM Tris-hydrochloric acid suffer solution, pH 6.8, 120 mM 2-mercaptoethanol, 2% SDS solution, 0.025% bromophenol blue, and 20% glycerol) and the resulting solution was heated at 95°C for 3 minutes and then subjected to SDS-polyacrylamide gel electrophoresis. The molecular weight of

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the translation product was determined by carrying out the autoradiography.

(7) Expression in COS7

Escherichia coli bearing a vector expressing the protein 5 of the invention was infected with helper phage M13K07, and single-stranded phage particles were obtained according to the method as stated above. Using the thus obtained phages, each expression vecotr was introduced into simian-kidney-origin culture cells COS7 in the manner as stated above. 10 incubation at 37 °C for 2 days in the presence of 5 % CO2, further incubation was carried out in a medium containing $[^{35}S]$ cysteine or $[^{35}S]$ methionine for 1 hour. The cells were collected, dissolved and then subjected to SDS-PAGE whereby a band corresponding to the expression product of each protein 15 which is not present in COS7 cells was revealed. In Table 3, the molecular weight of each expression product is shown.

Table 3

HP Number	Supernatant of culture	Membrane fraction
	(kDa)	(kDa)
HP01263	50	-
HP01299	-	30
HP01526	-	22
HP10230	-	24
HP10408	-	7
HP10415	· -	45
HP10424		14
HP10429	-	27
HP10432	-	17
HP10480	-	22

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(8) Clone Examples

<HP01263> (Sequence Number 1, 19, 37)

Determination of the whole base sequence for the cDNA insert of clone HP01263 obtained from the human liver cDNA 5 libraries revealed the structure consisting of a 5'-nontranslation region of 36 bp, an ORF of 1149 bp, and a 3'-nontranslation region of 316 bp. The ORF codes for a protein consisting of 382 amino acid residues with one transmembrane domain at the N-terminal. Figure 2 depicts the hydrophobicity 10 /hydrophilicity profile of the present protein obtained by the The in vitro translation resulted in Kyte-Doolittle method. formation of a translation product of 42 kDa, which is almost consistent with the molecular weight of 42,054 as predicted On expression in COS cells, an expression from the ORF. 50 kDa was observed in the culture 15 product of about supernatant. Therefore, said protein can be understood to be a secreted protein. Application of the rule (-3, -1) as a method for anticipation of a cutting site in a secretion signal sequence suggested that the mature protein would start from 20 methionine at 19 position.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the human α -2-HS-glycoprotein (SWISS-PROT Accession No. P02765). Table 4 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the human α -2-HS-glycoprotein (GP). represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the

protein of the present invention. The both proteins possessed a homology of 25.5%. The cysteine position is reserved and this region is analogous to that in cystatins (thiol proteinase inhibitors). There are observed other analogy with histidine-rich glycoprotein (P04196, 30.9%/194 amino acid residues), kininogen (P01045, 24.1%/261 amino acid residues), tyrosine kinase inhibitor (A32827, 24.4%/291 amino acid residues), and so on.

Table 4

10 HP MGLLLPLALCILVLCCGAMSPPQLALNPSALLSR--GCNDSDVLAVAGFALRDINKDRKD .*.** * . ..*. * .*.*... ..* *. **.. GP MKSLVLLLCLAQLWGCHSAPHGPGLIYRQPNCDDPETEEAALVAIDYINQNLPW HP GYVLRLNRVNDAQEYRRGGLGSLFYLTLDVLETDCHVLRKKAWQDCGMRIFFE-SVYGQC 15 GP GYKHTLNQIDEVKVWPQQPSGELFEIEIDTLETTCHVLDPTPVARCSVRQLKEHAVEGDC HP K-AIFYMNNPSRVLYLAAYNCTLRPVSKKKIYMTCPDCPSSIPTDSSNHQVLEAATESLA GP DFQLLKLDGKFSVVY---AKCDSSPDSAEDVRKVCQDCPLLAPLN--DTRVVHAAKAALA 20 HP KYNNENTSKQYSLFKVTRASSQWVVGPSYFVEYLIKESPC---TKSQASSCSLQSSDSVP . ** .**. * . ..*..*.*..*. * ...** GP AFNAQNNGSNFQLEEISRAQLV-PLPPSTYVEFTVSGTDCVAKEATEAAKCNLLAEKQY-HP VGLCKGSLTRTHWEKFVSVTCDFFESQAPATGSENSAVNQK-PTNLPKVEESQQKNTPPT *.**..*. . . *.***. *..*.*. ** 25 GP -GFCKATLSEKLGGAEVAVTCTVFQTQPVTSQPQPEGANEAVPTPVVDPDAPPSPPLGAP HP DSPSKAGPRGSVQYLPDLDDKNSQEKGPQEAFPVHLDLTTNPQGETLDISFLFLEPMEEK . *. ..*..* *. GP GLPPAGSPPDSHVLLAAPPGHQLHRAHYDLRHTFMGVVSLGSPSGEVSHPRKTRTVVQPS HP LVVLPFPKEKARTAECPGPAQNASPLVLPP 30 GP VGAAAGPVVPPCPGRIRHFKV

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Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. H57204), but it can not be assessed whether these ESTs with 5 partial sequences code for the same protein as the protein of the present invention. Hereupon, most of ESTs matching with the present cDNA are available from liver cDNA libraries, whereby the present clone is considered to be expressed specifically in the liver.

The present protein, because of being a type-II membrane protein, is considered to exert its function as a receptor on the membrane surface with the C-terminal side exposed outside the cells or after undergoing a processing followed by being excreted in the serum. The present protein, because of bearing 15 a cystatin-like domain, is considered to possess a proteinaseinhibitor activity as well as many physiological activities in the same manner as for other members of this family. In addition, the present protein, because of being expressed specifically in liver cells, is considered to play a 20 significant role for maintaining the liver function.

<HP01299> (Sequence Number 2, 20, 38)

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Determination of the whole base sequence for the cDNA insert of clone HP01299 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-non-25 translation region of 110 bp, an ORF of 954 bp, and a 3'-nontranslation region of 285 bp. The ORF codes for a protein consisting of 317 amino acid residues with two or more depicts the transmembrane 3 domains. Figure hydrophobicity/hydrophilicity profile of the present protein

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obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 32 kDa that was almost consistent with the molecular weight of 35,965 predicted from the ORF.

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The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the rat retinol dehydrogenase (NBRF Accession No. A55884). Table 5 indicates the comparison of the amino acid sequences between the human protein of the present invention 10 (HP) and the rat retinol dehydrogenase (RN). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and. represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 65.3% 15 among the entire regions.

Table 5

		•
	HP	MWLYLAAFVGLYYLLHWYRERQVVSHLQDKYVFITGCDSGFGNLLARQLDARGLRVLAAC
5		***** *.***. *****.*****************
	RN	${\tt MWLYLLALVGLWNLLRLFRERKVVSHLQDKYVFITGCDSGFGNLLARQLDRRGMRVLAAC}$
	HP	LTEKGAEQLRGQTSDRLETVTLDVTKMESIAAATQWVKEHVGDRGLWGLVNNAGILTPIT

	RN	LTEKGAEQLRSKTSDRLETVILDVTKTESIVAATQWVKERVGNRGLWGLVNNAGISVPVG
10	HP	LCEWLNTEDSMNMLKVNLIGVIQVTLSMLPLVRRARGRIVNVSSILGRVAFFVGGYCVSK
		****.***.***.***.***.**.**.***
	RN	PNEWMRKKDFASVLDVNLLGVIEVTLNMLPLVRKARGRVVNIASTMGRMSLVGGGYCISK
	ĦР	YGVEAFSDILRREIQHFGVKISIVEPGYFRTGMTNMTQSLERMKQSWKEAPKHIKETYGQ
		******* ******** *.*****. *
15	RN	YGVEAFSDSLRRELTYFGVKVAIIEPGGFKTNVTNMERLSDNLKKLWDQTTEEVKEIYGE
	ĦP	QYFDALYNIMKEGLLNCSTNLNLVTDCMEHALTSVHPRTRYSAGWDAKFFFIPLSYLPTS
		* ****.******* ******.*****
	RN	KFQDSYMKAMESLVNTCSGDLSLVTDCMEHALTSCHPRTRYSPGWDAKFFYLPMSYLPTF
	HP	LADYILTRSWPKPAQAV
20		*.* ***.*.
	RN	LSDAVIHWGSVKPARAL

Furthermore, the search of GenBank using the base sequence
25 of the present cDNA revealed that there existed some ESTs
possessing the homology of 90% or more (for example, Accession
No. R35197), but any of them was shorter than the present cDNA
and did not contain the initiation codon.

The rat retinol dehydrogenase has been found as a 30 microsomal membrane protein participating in the retinoic acid

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biosynthesis in the liver [Chai, X. et al., J. Biol. Chem. 270: 28408-28412 (1995)]. Accordingly, its homologue, the protein of the present invention, is considered to possess a similar function and can be utilized for diagnosis and treatment of 5 diseases caused by the abnormality of this protein.

<HP01347> (Sequence Number 3, 21, 39)

Determination of the whole base sequence for the cDNA insert of clone HP01347 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-non-10 translation region of 24 bp, an ORF of 891 bp, and a 3'-nontranslation region of 728 bp. The ORF codes for a protein consisting of 296 amino acid residues with one transmembrane depicts N-terminal. Figure the domain hydrophobicity/hydrophilicity profile of the present protein 15 obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified and the urokinase activity was detected on the membrane surface, upon transduction into the COS7 cells of an expression vector 20 in which a HindIII-SacI fragment (treated with the mung-bean nuclease) containing a cDNA fragment encoding the N-terminal 73 amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. Therefore, the present protein is considered to be a type-II membrane protein. The in vitro 25 translation resulted in the formation of a translation product of 33 kDa that was almost consistent with the molecular weight of 33,527 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was

analogous to the human HIV envelope glycoprotein gp120-binding C-type lectin (GenBank Accession No. M98457). Table 6 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the human HIV envelope glycoprotein gp120-binding C-type lectin (CL). represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 85.6% among 284 amino acid residues. There is observed at the downstream of the transmembrane domain a sequence with seven repetition of Ile-Tyr-Gln-Xaa-Leu-Thr-Xaa-Leu-Lys-Ala-Ala-Val-Gly-Glu-Leu-Xaa-Xaa-Xaa-Ser-Lys-Xaa-Gln-Xaa.

Table 6

	HP	MSDSKEPRVQQLGLLGCLGHGALVLQLLSFMLLAGVLVAI
		******* ***** ***** ****
5	CL	${\tt MSDSKEPRLQQLGLLEEEQLRGLGFRQTRGYKSLAGCLGHGPLVLQLLSFTLLAGL}$
	HP	${\tt LVQVSKVPSSLSQEQSEQDAIYQNLTQLKAAVGELSEKSKLQEIYQELTQLKAAVGELPE}$

	CL	LVQVSKVPSSISQEQSRQDAIYQNLTQLKAAVGELSEKSKLQEIYQELTQLKAAVGELPE
	HP	KSKLQEIYQELTRLKAAVGELPEKSKLQEIYQELTRLKAAVGELPEKSKLQEIYQELTRL
10		******************
	CL	KSKLQEIYQELTRLKAAVGELPEKSKLQEIYQELTWLKAAVGELPEKSKMQEIYQELTRL
	HP	KAAVGELPEKSKLQEIYQELTELKAAVGELPEKSKLQEIYQELTQLKAAVGELPDQSKQQ
		*********** ****** ******* ******* *****
	CL	KAAVGELPEKSKQQEIYQELTRLKAAVGELPEKSKQQEIYQELTRLKAAVGELPEKSKQQ
15	HP	QIYQELTDLKTAFERLCRHCPKDWTFFQGNCYFMSNSQRNWHDSVTACQEVRAQLVVIKT

	CL	EIYQELTQLKAAVERLCHPCPWEWTFFQGNCYFMSNSQRNWHDSITACKEVGAQLVVIKS
	HP	AEEQLPAVLEQWRTQQ
		**** *. *
20	CL	AEEQNFLQLQSSRSNRFTWMGLSDLNQEGTWQWVDGSPLLPSFKQYWNRGEPNNVGEEDC

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. H90360), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

The present protein, because of being a type-II membrane 30 protein, is considered to exert its function as a receptor on

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the membrane surface with the C-terminal side exposed outside the cells or after undergoing a processing followed by being excreted in the serum. Hereupon, the human HIV envelope glycoprotein gp120-binding C-type lectin that is highly 5 homologous with the present protein has been found as a CD4independent HIV receptor [Curtis, B. M. et al., Proc. Natl. Acad. Sci. USA 89: 8356-8360 (1992)].

<HP01440> (Sequence Number 4, 22, 40)

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Determination of the whole base sequence for the cDNA 10 insert of clone HP01440 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 37 bp, an ORF of 594 bp, and a 3'-nontranslation region of 98 bp. The ORF codes for a protein consisting of 197 amino acid residues with four transmembrane 15 domains. Figure 5 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 21 kDa that was almost consistent with the molecular weight of 20,822 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the human tumor-associated antigen L6 (SWISS-PROT Accession No. P30408). Table 7 indicates the comparison of the amino acid sequences between the human protein of the present 25 invention (HP) and the human tumor-associated antigen L6 (L6). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed

a homology of 47.0% among the entire regions.

Table 7

- Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more and also containing the initiation codon (for example, Accession No. T55097), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.
- 25 The human tumor-associated antigen L6 is a member of a membrane antigen TM4 superfamily proteins which are expressed in large quantities on the surface of human tumor cells [Marken, J. S. et al., Proc. Natl. Acad. Sci. USA 89: 3503-3507 (1992)]. Since these membrane antigens are expressed specifically on some specified cells or cancer cells,

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antibodies against these antigens, if constructed, are useful for a variety of diagnoses and as carriers for the drug delivery. In addition, the cells in which genes of these membrane antigens are transduced and the membrane antigens are expressed are applicable for detection of the corresponding ligands and so on.

<HP01526> (Sequence Number 5, 23, 41)

Determination of the whole base sequence for the cDNA insert of clone HP01526 obtained from the human stomach cancer 10 cDNA libraries revealed the structure consisting of a 5'-non-translation region of 83 bp, an ORF of 666 bp, and a 3'-non-translation region of 573 bp. The ORF codes for a protein consisting of 221 amino acid residues with a hydrophobic region of putative six transmembrane domains. Figure 6 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 23 kDa that was almost consistent with the molecular weight of 25,030 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the mouse interstitial cell protein (GenBank Accession No. X96618). Table 8 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the mouse interstitial cell protein (MM). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present inventions possessed

a homology of 79.6% among the entire regions.

Table 8

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more and also containing the initiation codon (for example, Accession No. H02682), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

25 The mouse interstitial cell protein has been cloned as a membrane protein that is expressed with highly increasing in interstitial cells stimulated by a cytokine [Tagoh, H. et al., Biochem. Biophys. Res. Commun. 221: 744-749 (1996)]. Since these membrane proteins are expressed specifically on some specified cells and cancer cells, antibodies against these

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proteins, if constructed, are useful for a variety of diagnoses and as carriers for the drug delivery. In addition, the cells in which genes of these membrane antigens are transduced and the membrane antigens are expressed are applicable for detection of the corresponding ligands and so on.

<HP10230> (Sequence Number 6, 24, 42)

Determination of the whole base sequence for the cDNA insert of clone HP10230 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-10 translation region of 190 bp, an ORF of 756 bp, and a 3'-nontranslation region of 2099 bp. The ORF codes for a protein consisting of 251 amino acid residues with at least one depicts the Figure 7 domain. transmembrane hydrophobicity/hydrophilicity profile of the present protein 15 obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 30 kDa that was almost consistent with the molecular weight of 28,800 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the nematode hypothetical protein F25D7.1 (GenBank Accession No. 278418). Table 9 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the nematode hypothetical protein F25D7.1 (CE). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 49.8% among the entire regions.

Table 9

HS MSDIGDWFRSIPAITRYWFAATVAVPLVGKLGLISPAYLFL-WPEAFLYRFQIWRPITAT CE MDLENFLLGIPIVTRYWFLASTIIPLLGRFGFINVQWMFLQW-DLVVNKFQFWRPLTAL 5 HS FYFPVGPGTGFLYLVNLYFLYQYSTRLETGAFDGRPADYLFMLLFNW-ICIVITGLAMDM CE IYYPVTPOTGFHWLMMCYFLYNYSKALESETYRGRSADYLFMLIFNWFFCSGLC-MALDI HS QLLMIPLIMSVLYVWAQLNRDMIVSFWFGTRFKACYLPWVILGFNYIIGGSVINELIGNL .*. *...***** *.*.* ****** ** * *****. *** .. *. .***.* 10 CE YFLLEPMVISVLYVWCQVNKDTIVSFWFGMRFPARYLPWVLWGFNAVLRGGGTNELVGIL HS VGHLYFFLMFRYPMDLGGRNFLSTPQFLYRWLPSRRGGVSGFGVPPASMRRAADQNGGGG *** ***. ..** . * ...***.* .*. **. * CE VGHAYFFVALKYPDEYGV-DLISTPEFLHRLIPDEDGGIHG---QDGNIRGARQQPRG--15 HS RHNW--GQGFRLGDQ * * * * *** CE -HQWPGGVGARLGGN

20 Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more and also containing the initiation codon (for example, Accession No. W01493), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

<HP10389> (Sequence Number 7, 25, 43)

Determination of the whole base sequence for the cDNA insert of clone HP10389 obtained from the human epidermoid carcinoma cell line KBc cDNA libraries revealed the structure consisting of a 5'-non-translation region of 62 bp, an ORF of

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321 bp, and a 3'-non-translation region of 270 bp. The ORF codes for a protein consisting of 106 amino acid residues with a hydrophobic region of putative two transmembrane domains. Figure 8 depicts the hydrophobicity/hydrophilicity profile of 5 the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 12 kDa that was almost consistent with the molecular weight of 11,528 predicted from the ORF.

The search of the protein data base using the amino acid 10 sequence of the present protein revealed that the protein was not analogous to any of known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. H70816), but many sequences 15 were not distinct and the same ORF as that in the present cDNA was not identified.

<HP10408> (Sequence Number 8, 26, 44)

Determination of the whole base sequence for the cDNA insert of clone HP10408 obtained from the human stomach cancer 20 cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 74 bp, an ORF of 237 bp, and a 3'-nontranslation region of 128 bp. The ORF codes for a protein consisting of 78 amino acid residues with a putative signal sequence at the N-terminal as well as a sequence of one 25 putative interior transmembrane domain. Figure 9 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified WO 98/55508 PCT/JP98/02445

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upon transduction into the COS7 cells of an expression vector in which a HindIII-BglII fragment (after the Klenow treatment) containing a cDNA fragment encoding the N-terminal 70 amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. The in vitro translation resulted in the formation of a translation product of 9 kDa that was almost consistent with the molecular weight of 8,396 predicted from the ORF.

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. T94049), but they were shorter than the present cDNA and any molecule containing the initiation codon was not identified.

15 <HP10412> (Sequence Number 9, 27, 45)

Determination of the whole base sequence for the cDNA insert of clone HP10412 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 55 bp, an ORF of 945 bp, and a 3'-non-20 translation region of 131 bp. The ORF codes for a protein consisting of 314 amino acid residues with one transmembrane the 10 depicts Figure N-terminal. at the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that 25 the present protein remained in the membrane from the observation that the urokinase secretion was not identified upon transduction into the COS7 cells of an expression vector in which a HindIII-ApaI fragment (treated with mung-bean nuclease) containing a cDNA fragment encoding the N-terminal 65 amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. The in vitro translation resulted in the formation of a translation product of 44 kDa that was somewhat larger than the molecular weight of 35,610 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the nematode hypothetical protein of 28.5 kDa (SWISS-PROT Accession No. P34623). Table 10 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the nematode hypothetical protein of 28.5 kDa (CE). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 42.8% in the C-terminal region of 243 amino acid residues.

Table 10

HP MVAPVWYLVAAALLVGFILFLTRSRGRAASAGQEPLHNEELAGAGRVAQPGPLEPEEPRA 5 HP GGRPRRRDLGSRLQAQRRAQRVAWAEA--DENEEEAVILAQEEEGVEKPAETHLSGKIG * .*.**..**.*. *** MRRNARRRVNRDEQEDGFVNHMMNDGEDVEDLDGGAEQFEYDEDGKKIG CE HP AKKLRKLEEKQARKAQREAEEAEREERKRLESQREAEWKKEEERLRLEEEQKEEEE--RK .* **..*... ** * ****** *..* * *..*** . *...*.** 10 CE KRKAAKLQAKEEKRQMREYEVREREERKRREEER--EKKRDEERAKEEADEKAEEERLRK HP AREEQAQREHEEYLKLKEAFVVEEEGVGETMTEEQSQSFLTEFINYIKQSKVVLLEDLAS CE EREKERKEHEEYLAMKASFAIEEEG-TDAIEGEEAENLIRDFVDYVKTNKVVNIDELSS HP QVGLRTQDTINRIQDLLAEGTITGVIDDRGKFIYITPEELAAVANFIRQRGRVSIAELAQ 15 CE HFGLKSEDAVNRLQHFIEEGLVQGVMDDRGKFIYISDEEFAAVAKFINQRGRVSIHEIAE HP ASNSLIAWGRESPAQAPA .**.** . *.*. CE QSNRLIRLETPSAAE 20

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. T09311), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10413> (Sequence Number 10, 28, 46)

Determination of the whole base sequence for the cDNA 30 insert of clone HP10413 obtained from the human stomach cancer

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cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 78 bp, an ORF of 588 bp, and a 3'-nontranslation region of 1209 bp. The ORF codes for a protein consisting of 195 amino acid residues with one transmembrane depicts the N-terminal. Figure 11 5 domain at hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified 10 upon transduction into the COS7 cells of an expression vector in which a HindIII-PmaCI fragment containing a cDNA fragment encoding the N-terminal 65 amino acid residues in the present protein was inserted at the HindIII-PmaCI site of pSSD3. The in vitro translation resulted in the formation of a translation 15 product of 28 kDa that was somewhat larger than the molecular weight of 21,671 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the swine steroidal membrane-binding protein (GenBank Accession No. X99714). Table 11 indicates the comparison of the amino acid sequences between the human protein of the present invention (HP) and the swine steroidal membrane-binding protein (SS). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 96.4% among the entire regions.

Table 11

	ĦР	${\tt MAAEDVVATGADPSDLESGGLLHEIFTSPLNLLLLGLCIFLLYKIVRGDQPAASGDSDDD}$
		****** ******* ** ****************
5	ss	MAAEDVAATGADPSELEGGGLLHEIFTSPLNLLLLGLCIFLLYKIVRGDQPAAS-DSDDD
	нР	EPPPLPRLKRRDFTPAELRRFDGVQDPR1LMAINGKVFDVTKGRKFYGPEGPYGVFAGRD

	SS	EPPPLPRLKRRDFTPAELRRFDGVQDPRILMAINGKVFDVTKGRKFYGPEGPYGVFAGRD
•	ĦР	ASRGLATFCLDKEALKDEYDDLSDLTAAQQETLSDWESQFTFKYHHVGKLLKEGEEPTVY
10		******************
	SS	ASRGLATFCLDKEALKDEYDDLSDLTPAQQETLNDWDSQFTFKYHHVGKLLKEGEEPTVY
	ĦР	SDEEEPKDESARKND

	SS	SDEEEPKDESARKND
15		

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA021062), but many sequences were not distinct and the same ORF as that in the present cDNA was not identified.

<HP10415> (Sequence Number 11, 29, 47)

Determination of the whole base sequence for the cDNA insert of clone HP10415 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 71 bp, an ORF of 1389 bp, and a 3'-non-translation region of 103 bp. The ORF codes for a protein consisting of 462 amino acid residues with one transmembrane domain at the N-terminal. Figure 12 depicts the hydrophobicity/hydrophilicity profile of the present protein

obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 48 kDa that was somewhat smaller than the molecular weight of 52,458 predicted from the ORF.

5 The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the cytochrome P450 as exemplified by the simian cytochrome P450IIIA8 (SWISS-PROT Accession No. P33268). Table 12 indicates the comparison of the amino acid sequences between 10 the human protein of the present invention (HP) and the simian cytochrome P450IIIA8 (CP). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present invention. The both proteins possessed a homology of 21.3% among the entire regions.

Table 12

	HP	MLDFAIFAVTFLLALVGAVLYLYPASRQAAGIPGITPTEEKDGNLPDIVN-SGSLHEF
		.******** .* .* .* .* .* .*
5	CP	MDLIPDLAVETWLLLAVTLVLLYLYGTHSHGLFKKLGIPGPTPLPLLGNILSYRKGFWTF
	HP	LVNLHERYGPVVSFWFGRRLVVSLGTVDVLKQHINPNKTLDPFETMLK-SLLRYQSGGGS
		** * .*. **. * * * *
	CP	DMECYKKYGKVWGFYDGRQPVLAITDPNMIK-TVLVKECYSVFTNRRPFGPVGFMKNAIS
	HP	VSENHMRKKLYENGVTDSLKSNFALLLKLSEELLDKWLSYPET-QHVPLSQHMLGF
10		**. *** * ****.*
	CP	IAEDEEWKRIRSLLSPTFTSGKLKEMVPIIAKYGDVLVRNLRREAETGKPVTLKDVFGAY
	HP	AMKSVTQMVMGSTF-EDDQEVIRFQKNHGTVWSEIGKGFLDGSLDKNM
		.** .* * *. *
	СP	SMDVITSTSFGVNIDSLNNPQDPFVENTKKLLRFDFLDPFFLSITIFPFIIPILEVLNIS
15	HP	TRKKQYEDALMQ-LESVLRNIIKE-RKGR-NFSQHIFIDSLVQGNLNDQQILEDS
		*
	CP	IFPREVTSFLRKSVKRIKESRLKDTQKHRVDFLQLMIDSQNSKETESHKALSDLELVAQS
	HP	MIFSLASCIITAKLCTWAICFLTTSEEVQKKLYEEINQVF-GNGPVTPEKIEQLRYCQHV
		** .** *. *. * .** .** . * * * * . * * * * . * * * * * * * * * * * * *
20	CP	IIFIFAGYETTSSVLSFIIYELATHPDVQQKLQEEIDTVLPNKAPPTYDTVLQMEYLDMV
	HP	LCETVRTAKLTPVSAQLQDIEGKIDRFIIPRETLVLYALGVVLQDPNTWPSPHKFDPDRF
		. **.*
	HP	VNETLRIFPIAMRLERVCKKDVEINGIFIPKGVVVMIPSYALHHDPKYWPEPEKFLPERF
	HP	DDELVMKTFSSLGFSGTQECPELRFAYMVTTVLLSVLVKRLHLLSVEGQVIETKYE
25		.* ***** * * *
	CP	SKKNNDNIDPYIYTPFG-SGPRNCIGMRFALMNMKLAIIRVLQNFSFKPCKETQIPLKLR
	HP	LVTSSREEAWITVSKRY
		*
	CP	LGGLLQTEKPIVLKIESRDGTVSGA

Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs

possessing the homology of 90% or more (for example, Accession No. AA381169), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

The cytochrome P450 participates in the drug metabolism and can be utilized as a catalyst in organic synthesis reactions such as oxidation and so on.

<HP10419> (Sequence Number 12, 30, 48)

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Determination of the whole base sequence for the cDNA insert of clone HP10419 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 170 bp, an ORF of 744 bp, and a 3'-non-translation region of 1116 bp. The ORF codes for a protein consisting of 247 amino acid residues with a hydrophobic region of putative seven transmembrane domains. Figure 13 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method.

The search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA340663), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10424> (Sequence Number 13, 31, 49)

Determination of the whole base sequence for the cDNA insert of clone HP10424 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-non-translation region of 97 bp, an ORF of 342 bp, and a 3'-non-translation region of 54 bp. The ORF codes for a protein

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consisting of 113 amino acid residues with one transmembrane depicts Figure 14 the N-terminal. at domain hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that 5 the present protein remained in the membrane from the observation that the urokinase secretion was not identified upon transduction into the COS7 cells of an expression vector in which a HindIII-AccI fragment (after the Klenow treatment) containing a cDNA fragment encoding the N-terminal 58 amino 10 acid residues in the present protein was inserted at the HindIII-SmaI site of pSSD3. The in vitro translation resulted in the formation of a translation product of 14 kDa that was somewhat larger than the molecular weight of 12,784 predicted from the ORF.

of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA401979), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10428> (Sequence Number 14, 32, 50)

Determination of the whole base sequence for the cDNA insert of clone HP10428 obtained from the human epidermoid carcinoma cell line KBc cDNA libraries revealed the structure consisting of a 5'-non-translation region of 287 bp, an ORF of 1098 bp, and a 3'-non-translation region of 659 bp. The ORF codes for a protein consisting of 365 amino acid residues with a hydrophobic region of putative nine transmembrane domains. Figure 15 depicts the hydrophobicity/hydrophilicity profile of

the present protein obtained by the Kyte-Doolittle method. The result of the in vitro translation did not reveal the formation of distinct bands and only revealed the formation of smeary bands at the high-molecular-weight position.

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The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was analogous to the baker's yeast hypothetical membrane protein YML038c (NBRF Accession No. S49741). Table 13 indicates the comparison of the amino acid sequences between the human 10 protein of the present invention (HP) and the baker's yeast hypothetical membrane protein YML038c (SC). - represents a gap, * represents an amino acid residue identical to that in the protein of the present invention, and . represents an amino acid residue analogous to that in the protein of the present 15 invention. The both proteins possessed a homology of 26.3% among the N-terminal region of 281 amino acid residues.

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Table 13

	HP	MGRWALDVAFLWKAVLTLGLVL-LYYCFSIGITFYNKWLTKSFHFPLFMTMLHLA
		. *.* *.*
5	sc	MNRTVFLAFVFGWYFCS-IALSIYNRWMFDPKDGLGIGYPVLVTTFHQA
	HP	VIFLFSALSRALVQCSSHRARVVLSWADYLRRVAPTALATALDVGLSNWSFLYVTVS
		.. * ** * * * . * * * * * * * * * * *
	sc	TLWLLSGIYIKLRHKPVKNVLRKNNGFNWSFFLKFLLPTAVASAGDIGLSNVSFQYVPLT
	нр	LYTMTKSSAVLFILIFSLIFKLEELRAALVLVVLLIAGGLFMFTYKSTQ-FN
10		_***** *.*. ***** **** . ** . ** .
	sc	IYTIIKSSSIAFVLLFGCIFKLEKFHWKLALSVIIMFVGVALMVFKPSDSTSTKNDQALV
	HP	VEGFALVLGASFIGGIRWTLTQMLLQKAELGLQNPIDTMFHLQPLMFLGLFPLFAVFEGL
		. * *****.**. ***
	sc	IFGSFLVLASSCLSGLRWVYTQLMLRNNPIQTNTAAAVEES-DGALFTENEDNVDNEPVV
15	HP	HLSTSEKIFRFQDT-GLLLRVLGSLFLGGILAFGLGFSEFLLVSRTSSLTLSIAGIFKEV
		.** * *. ** *** ***
	sc	NLANNKMLENFGESKPHPIHTIHQLAPIMGITLLLTS-LLVEKPFPGIFS-SSIFRLD
	нР	CTLLLAAHLLGDQISLLNWLGFALCLSGISLHVALKALHSRGDGGPKALKGLGSSPDLEL
20	sc	TSNGGVGTETTVLSIVRGIVLLILPGFAVFLLTICEFSILEQTPVLTVSIVGIVKELLTV
	HP	LLRSSQREEGDNEEEEYFVAQGQQ
	sc	IFGIIILSERLSGFYNWLGMLIIMADVCYYNYFRYKQDLLQKYHSVSTQDNRNELKGFQD

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Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. AA018345), but it can not be assessed whether these ESTs

with partial sequences code for the same protein as the protein of the present invention.

<HP10429> (Sequence Number 15, 33, 51)

Determination of the whole base sequence for the cDNA 5 insert of clone HP10429 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 156 bp, an ORF of 681 bp, and a 3'-nontranslation region of 206 bp. The ORF codes for a protein consisting of 226 amino acid residues with four transmembrane 10 domains. Figure 16 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 25 kDa that was almost consistent with the molecular weight of 25,321 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or 20 more (for example, Accession No. AA315933), but it can not be assessed whether these ESTs with partial sequences code for the same protein as the protein of the present invention.

<HP10432> (Sequence Number 16, 34, 52)

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Determination of the whole base sequence for the cDNA 25 insert of clone HP10429 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 28 bp, an ORF of 390 bp, and a 3'-nontranslation region of 554 bp. The ORF codes for a protein consisting of 129 amino acid residues with a signal-like

sequence at the N-terminal and one interior transmembrane domain. Therefore, the present protein is considered to be a type-I membrane protein. Figure 17 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. T74424), but the same ORF as that in the present cDNA was not identified.

<HP10433> (Sequence Number 17, 35, 53)

Determination of the whole base sequence for the cDNA insert of clone HP10433 obtained from the human liver cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 72 bp, an ORF of 492 bp, and a 3'-nontranslation region of 131 bp. The ORF codes for a protein consisting of 163 amino acid residues with one transmembrane 18 depicts N-terminal. Figure 20 domain the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. It was indicated that the present protein remained in the membrane from the observation that the urokinase secretion was not identified 25 upon transduction into the COS7 cells of an expression vector in which a HindIII-Eco81I fragment (treated with the mung-bean nuclease) containing a CDNA fragment encoding the N-terminal 137 amino acid residues in the present protein was inserted at the HindIII-EcoRV site of pSSD3. Therefore, the present protein

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is considered to be a type-II membrane protein. The in vitro translation resulted in the formation of a translation product of 21 kDa that was almost consistent with the molecular weight of 18,617 predicted from the ORF.

The search of the protein data base using the amino acid sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or 10 more (for example, Accession No. H84693), but many sequences are not distinct and the same ORF as that in the present cDNA was not identified.

<HP10480> (Sequence Number 18, 36, 54)

Determination of the whole base sequence for the cDNA 15 insert of clone HP10480 obtained from the human stomach cancer cDNA libraries revealed the structure consisting of a 5'-nontranslation region of 79 bp, an ORF of 582 bp, and a 3'-nontranslation region of 1253 bp. The ORF codes for a protein consisting of 193 amino acid residues with four transmembrane 20 domains. Figure 19 depicts the hydrophobicity/hydrophilicity profile of the present protein obtained by the Kyte-Doolittle method. The in vitro translation resulted in the formation of a translation product of 23 kDa that was somewhat larger than the molecular weight of 21,445 predicted from the ORF.

The search of the protein data base using the amino acid 25 sequence of the present protein revealed that the protein was not analogous to any known proteins. Furthermore, the search of GenBank using the base sequence of the present cDNA revealed that there existed some ESTs possessing the homology of 90% or more (for example, Accession No. W93606), but many sequences are not distinct and the same ORF as that in the present cDNA was not identified.

The present invention provides human proteins having transmembrane domains and cDNAs encoding said proteins. All of the proteins of the present invention are putative proteins controlling the proliferation and differentiation of the cells, because said proteins exist on the cell membrane. Therefore, the proteins of the present invention can be used as pharmaceuticals or as antigens for preparing antibodies against said proteins. Furthermore, said DNAs can be used for the expression of large amounts of said proteins. The cells expressing large amounts of membrane proteins with transfection of these membrane protein genes can be applied to the detection of the corresponding ligands, the screening of novel low-molecular medicines, and so on.

In addition to the activities and uses described above, the polynucleotides and proteins of the present invention may exhibit one or more of the uses or biological activities (including those associated with assays cited herein) identified below. Uses or activities described for proteins of the present invention may be provided by administration or use of such proteins or by administration or use of polynucleotides encoding such proteins (such as, for example, in gene therapies or vectors suitable for introduction of DNA).

Research Uses and Utilities

The polynucleotides provided by the present invention can be used by the research community for various purposes. The polynucleotides can be used to express recombinant protein for

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analysis, characterization or therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as 5 molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to identify chromosomes or to map related gene positions; to compare with endogenous DNA sequences in patients to identify potential genetic disorders; as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR primers for genetic fingerprinting; as a probe to "subtract-out" known the process of discovering other in sequences polynucleotides; for selecting and making oligomers for attachment to a "gene chip" or other support, including for 15 examination of expression patterns; to raise anti-protein antibodiesusing DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or potentially binds to another protein (such as, for example, in 20 a receptor-ligand interaction), the polynucleotide can also be used in interaction trap assays (such as, for example, that described in Gyuris et al., Cell 75:791-803 (1993)) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

25 The proteins provided by the present invention can similarly be used in assay to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in

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assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of 15 being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art. References disclosing such methods include without limitation "Molecular Cloning: A 20 Laboratory Manual", 2d ed., Cold Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology: Guide to Molecular Cloning Techniques", Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

25 <u>Nutritional Uses</u>

Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a nitrogen source

and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, 5 pills, solutions, suspensions or capsules. In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

Cytokine and Cell Proliferation/Differentiation

Activity 10

A protein of the present invention may exhibit cytokine, cell proliferation (either inducing or inhibiting) or cell differentiation (either inducing or inhibiting) activity or may other cytokines in certain induce production of Many protein factors discovered to date, 15 populations. including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein of the present invention 20 is evidenced by any one of a number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B9, B9/11, BaF3, MC9/G, M+ (preB M+), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7e and CMK.

The activity of a protein of the invention may, among 25 other means, be measured by the following methods:

Assays for T-cell or thymocyte proliferation include without limitation those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H.

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Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Takai et al., J. Immunol. 137:3494-3500, 1986; Bertagnolli et al., J. Immunol. 145:1706-1712, 1990; Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Bertagnolli, et al., J. Immunol. 149:3778-3783, 1992; Bowman et al., J. Immunol. 152: 1756-1761, 1994.

Assays for cytokine production and/or proliferation of spleen cells, lymph node cells or thymocytes include, without limitation, those described in: Po lyclonal T cell stimulation, Kruisbeek, A.M. and Shevach, E.M. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 3.12.1-3.12.14, John Wiley and Sons, Toronto. 1994; and Measurement of mouse and human Interferon γ, Schreiber, R.D. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.8.1-6.8.8, John Wiley and Sons, Toronto. 1994.

of differentiation proliferation and for Assays include, hematopoietic and lymphopoietic cells 20 limitation, those described in: Measurement of Human and Murine Interleukin 2 and Interleukin 4, Bottomly, K., Davis, L.S. and Lipsky, P.E. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.3.1-6.3.12, John Wiley and Sons, Toronto. 1991; deVries et al., J. Exp. Med. 173:1205-1211, 25 1991; Moreau et al., Nature 336:690-692, 1988; Greenberger et al., Proc. Natl. Acad. Sci. U.S.A. 80:2931-2938, 1983; Measurement of mouse and human interleukin 6 -Nordan, R. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.6.1-6.6.5, John Wiley and Sons, Toronto. 1991; Smith et 61

al., Proc. Natl. Acad. Sci. U.S.A. 83:1857-1861, 1986;
Measurement of human Interleukin 11 - Bennett, F., Giannotti,
J., Clark, S.C. and Turner, K. J. In Current Protocols in
Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 6.15.1 John Wiley
and Sons, Toronto. 1991; Measurement of mouse and human
Interleukin 9 - Ciarletta, A., Giannotti, J., Clark, S.C. and
Turner, K.J. In Current Protocols in Immunology. J.E.e.a.
Coligan eds. Vol 1 pp. 6.13.1, John Wiley and Sons, Toronto.
1991.

Assays for T-cell clone responses to antigens (which will 10 identify, among others, proteins that affect APC-T cell interactions as well as direct T-cell effects by measuring proliferation and cytokine production) include, without described in: Current Protocols limitation, those 15 Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function; Chapter 6, Cytokines and their cellular receptors; Chapter 7, Immunologic studies in Humans); 20 Weinberger et al., Proc. Natl. Acad. Sci. USA 77:6091-6095, 1980; Weinberger et al., Eur. J. Immun. 11:405-411, 1981; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988.

Immune Stimulating or Suppressing Activity

A protein of the present invention may also exhibit immune stimulating or immune suppressing activity, including without limitation the activities for which assays are described herein. A protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined

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immunodeficiency (SCID)), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial orfungal infections, or may result from autoimmune disorders. More specifically, infectious diseases causes by viral, bacterial, fungal or other infection may be treatable using a protein of the present invention, including infections by HIV, hepatitis viruses, herpesviruses, mycobacteria, Leishmania spp., malaria spp. and various fungal infections such as candidiasis. Of course, in this regard, a protein of the present invention may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Autoimmune disorders which may be treated using a protein 15 of the present invention include, for example, connective sclerosis, systemic multiple disease, erythematosus, rheumatoid arthritis, autoimmune pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, 20 insulin dependent diabetes mellitis, myasthenia gravis, graft-versus-host disease and autoimmune inflammatory eye disease. Such a protein of the present invention may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or 25 other respiratory problems. Other conditions, in which immune suppression is desired (including, for example, organ transplantation), may also be treatable using a protein of the present invention.

Using the proteins of the invention it may also be

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possible to immune responses, in a number of ways. Down regulation may be in the form of inhibiting or blocking an immune response already in progress or may involve preventing the induction of an immune response. The functions of 5 activated T cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or Immunosuppression of T cell responses is generally an both. active, non-antigen-specific, process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, 10 which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after exposure to the tolerizing agent has ceased. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure 15 to specific antigen in the absence of the tolerizing agent.

Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions (such as , for example, B7)), e.g., preventing high level lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue transplantation. Typically, in tissue transplants, rejection of the transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2

activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen blocking antibody), B7-1, B7-3) or prior (e.g., transplantation can lead to the binding of the molecule to the 5 natural ligand(s) on the immune cells without transmitting the corresponding costimulatory signal. Blocking B lymphocyte antigen function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an immunosuppressant. Moreover, the lack of costimulation may 10 also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigen-blocking reagents may avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the function of a combination of B lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4Ig fusion proteins in vivo as described in Lenschow et al., Science 257:789-792 (1992) and Turka et al., Proc. Natl. Acad. Sci USA, 89:11102-11105 (1992). In addition, murine models of GVHD (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function in vivo on the development of that disease.

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Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate activation of T cells that are reactive against self tissue and which promote the 5 production of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by disrupting receptor: ligand interactions of B lymphocyte 10 antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which may be involved in the disease process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead to long-term relief from the disease. 15 The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/lpr/lpr mice or NZB hybrid 20 mice, murine autoimmune collagen arthritis, diabetes mellitus in NOD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may be in the form of enhancing an existing immune response or eliciting an initial immune response. For example, enhancing an immune response through stimulating B

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lymphocyte antigen function may be useful in cases of viral In addition, systemic viral diseases such as infection. influenza, the commoncold, and encephalitis might be alleviated by the administration of stimulatory forms of B lymphocyte 5 antigens systemically.

Alternatively, anti-viral immune responses may be enhanced in an infected patient by removing T cells from the patient, costimulating the T cells in vitro with viral antigen-pulsed APCs either expressing a peptide of the present invention or 10 together with a stimulatory form of a soluble peptide of the present invention and reintroducing the in vitro activated T cells into the patient. Another method of enhancing anti-viral immune responses would be to isolate infected cells from a patient, transfect them with a nucleic acid encoding a protein 15 of the present invention as described herein such that the cells express all or a portion of the protein on their surface, and reintroduce the transfected cells into the patient. infected cells would now be capable of delivering a costimulatory signal to, and thereby activate, T cells in vivo.

In another application, up regulation or enhancement of antigen function (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least 25 one peptide of the present invention can be administered to a subject to overcome tumor-specific tolerance in the subject. If desired, the tumor cell can be transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected ex vivo with an expression

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vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression 5 of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection in vivo.

The presence of the peptide of the present invention having the activity of a B lymphocyte antigen(s) on the surface 10 of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack MHC class I or MHC class II molecules, or which fail to reexpress sufficient amounts of MHC class I or MHC class II 15 molecules, can be transfected with nucleic acid encoding all or a portion of (e.g., a cytoplasmic-domain truncated portion) of an MHC class I α chain protein and β_2 microglobulin protein or an MHC class II α chain protein and an MHC class II β chain protein to thereby express MHC class I or MHC class II proteins 20 on the cell surface. Expression of the appropriate class I or class II MHC in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which 25 blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a

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T cell mediated immune response in a human subject may be sufficient to overcome tumor-specific tolerance in the subject.

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The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for thymocyte or splenocyte cytotoxicity 5 include, without limitation, those described in: Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays 10 for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Herrmann et al., Proc. Natl. Acad. Sci. USA 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Herrmann et al., Proc. Natl. Acad. Sci. USA 15 78:2488-2492, 1981; Herrmann et al., J. Immunol. 128:1968-1974, 1982; Handa et al., J. Immunol. 135:1564-1572, 1985; Takai et al., J. Immunol. 137:3494-3500, 1986; Bowmanet al., J. Virology 61:1992-1998; Takai et al., J. Immunol. 140:508-512, 20 Bertagnolli et al., Cellular Immunology 133:327-341, 1991; Brown et al., J. Immunol. 153:3079-3092, 1994.

Assays for T-cell-dependent immunoglobulin responses and isotype switching (which will identify, among others, proteins that modulate T-cell dependent antibody responses and that affect Th1/Th2 profiles) include, without limitation, those described in: Maliszewski, J. Immunol. 144:3028-3033, 1990; and Assays for B cell function: In vitro antibody production, Mond, J.J. and Brunswick, M. In Current Protocols in Immunology. J.E.e.a. Coligan eds. Vol 1 pp. 3.8.1-3.8.16, John

Wiley and Sons, Toronto. 1994.

Mixed lymphocyte reaction (MLR) assays (which will identify, among others, proteins that generate predominantly Thl and CTL responses) include, without limitation, those described in: Current Protocols in Immunology, Ed by J. E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 3, In Vitro assays for Mouse Lymphocyte Function 3.1-3.19; Chapter 7, Immunologic studies in Humans); Takai et al., J. Immunol. 137:3494-3500, 1986; Takai et al., J. Immunol. 140:508-512, 1988; Bertagnolli et al., J. Immunol. 149:3778-3783, 1992.

Dendritic cell-dependent assays (which will identify, among others, proteins expressed by dendritic cells that 15 activate naive T-cells) include, without limitation, those described in: Guery et al., J. Immunol. 134:536-544, 1995; Inaba et al., Journal of Experimental Medicine 173:549-559, 1991; Macatonia et al., Journal of Immunology 154:5071-5079, 1995; Porgador et al., Journal of Experimental Medicine 182:255-260, 1995; Nair et al., Journal 67:4062-4069, 1993; Huang et al., Science 264:961-965, 1994; Macatonia et al., Journal of Experimental 169:1255-1264, 1989; Bhardwaj et al., Journal of Clinical Investigation 94:797-807, 1994; and Inaba et al., Journal of 25 Experimental Medicine 172:631-640, 1990.

Assays for lymphocyte survival/apoptosis (which will identify, among others, proteins that prevent apoptosis after superantigen induction and proteins that regulate lymphocyte homeostasis) include, without limitation, those described in:

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Darzynkiewicz et al., Cytometry 13:795-808, 1992; Gorczyca et al., Leukemia 7:659-670, 1993; Gorczyca et al., Cancer Research 53:1945-1951, 1993; Itoh et al., Cell 66:233-243, 1991; Zacharchuk, Journal of Immunology 145:4037-4045, 1990; Zamai et al., Cytometry 14:891-897, 1993; Gorczyca et al., International Journal of Oncology 1:639-648, 1992.

Assays for proteins that influence early steps of T-cell commitment and development include, without limitation, those described in: Antica et al., Blood 84:111-117, 1994; Fine et al., Cellular Immunology 155:111-122, 1994; Galy et al., Blood 85:2770-2778, 1995; Toki et al., Proc. Nat. Acad Sci. USA 88:7548-7551, 1991.

Hematopoiesis Regulating Activity

A protein of the present invention may be useful in 15 regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation 20 of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid cells; in supporting the 25 growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelo-suppression; in supporting the growth and proliferation of megakaryocytes and consequently

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of platelets thereby allowing prevention or treatment of various platelet disorders such as thrombocytopenia, and generally for use in place of or complimentary to platelet transfusions; and/or in supporting the growth and proliferation 5 of hematopoietic stem cells which are capable of maturing to any and all of the above-mentioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantation, including, without limitation, aplastic anemia and paroxysmal 10 nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in-vivo in conjunction with bone marrow ex-vivo (i.e., or progenitor with peripheral transplantation or transplantation (homologous or heterologous)) as normal cells 15 or genetically manipulated for gene therapy.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for proliferation and differentiation of various hematopoietic lines are cited above.

Assays for embryonic stem cell differentiation (which will identify, among others, proteins that influence embryonic differentiation hematopoiesis) include, without limitation, those described in: Johansson et al. Cellular Biology 15:141-151, 1995; Keller et al., Molecular and Cellular Biology 13:473-486, 1993; McClanahan et al., Blood 81:2903-2915, 1993.

Assays for stem cell survival and differentiation (which will identify, among others, proteins that regulate lympho-hematopoiesis) include, without limitation, those described in: Methylcellulose colony forming assays, Freshney,

M.G. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., Proc. Natl. Acad. Sci. USA 89:5907-5911, 1992; Primitive hematopoietic colony forming cells with high 5 proliferative potential, McNiece, I.K. and Briddell, R.A. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 23-39, Wiley-Liss, Inc., New York, NY. 1994; Neben et al., Experimental Hematology 22:353-359, 1994; Cobblestone area assay, In Culture Ploemacher, R.E. forming cell 10 Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 1-21, Wiley-Liss, Inc., New York, NY. 1994; Long term bone marrow cultures in the presence of stromal cells, Spooncer, E., Dexter, M. and Allen, T. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 163-179, Wiley-Liss, Inc., 15 New York, NY. 1994; Long term culture initiating cell assay, Sutherland, H.J. In Culture of Hematopoietic Cells. R.I. Freshney, et al. eds. Vol pp. 139-162, Wiley-Liss, Inc., New York, NY. 1994.

Tissue Growth Activity

- A protein of the present invention also may have utility 20 in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.
- A protein of the present invention, which induces 25 cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other Such a preparation employing a protein of the animals.

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invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. De novo bone formation induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

Another category of tissue regeneration activity that may be attributable to the protein of the present invention is tendon/ligament formation. A protein of the present invention, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue

formation induced by a composition of the present invention contributes to the repair of congenital, trauma induced, or other tendon or ligament defects of other origin, and is also useful in cosmetic plastic surgery for attachment or repair of The compositions of the present 5 tendons or ligaments. invention may provide an environment to attract tendon- or ligament-forming cells, stimulate growth of tendon- or ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of 10 tendon/ligament cells or progenitors ex vivo for return in vivo to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel syndrome and other tendon or ligament defects. The compositions may also include an appropriate matrix and/or 15 sequestering agent as a carrier as is well known in the art.

The protein of the present invention may also be useful for proliferation of neural cells and for regeneration of nerve and brain tissue, i.e. for the treatment of central and peripheral nervous system diseases and neuropathies, as well as 20 mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. specifically, a protein may be used in the treatment of diseases of the peripheral nervous system, such as peripheral and localized neuropathy injuries, peripheral 25 neuropathies, and central nervous system diseases, such as Huntington's disease, Alzheimer's, Parkinson's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome. Further conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders,

such as spinal cord disorders, head trauma and cerebrovascular diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

5 Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

It is expected that a protein of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to regenerate. A protein of the invention may also exhibit angiogenic activity.

A protein of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokine damage.

A protein of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for tissue generation activity include, without limitation, those described in: International Patent Publication No. W095/16035 (bone, cartilage, tendon); International Patent Publication No. W095/05846 (nerve, neuronal); International Patent Publication No. W091/07491 (skin, endothelium).

Assays for wound healing activity include, without limitation, those described in: Winter, Epidermal Wound Healing, pps. 71-112 (Maibach, HI and Rovee, DT, eds.), Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978).

Activin/Inhibin Activity

A protein of the present invention may also exhibit inhibin-related activities. Inhibins activinor 15 characterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins and are characterized by their ability to stimulate the release of follicle stimulating hormone (FSH). Thus, a protein of the present invention, alone or in heterodimers with a member of 20 the inhibin α family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, 25 as a homodimer or as a heterodimer with other protein subunits of the inhibin- β group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885. A protein of

the invention may also be useful for advancement of the onset of fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for activin/inhibin activity include, without limitation, those described in: Vale et al., Endocrinology 91:562-572, 1972; Ling et al., Nature 321:779-782, 1986; Vale et al., Nature 321:776-779, 1986; Mason et al., Nature 318:659-663, 1985; Forage et al., Proc. Natl. Acad. Sci. USA 83:3091-3095, 1986.

Chemotactic/Chemokinetic Activity

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A protein of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells. Chemotactic and chemokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action. Chemotactic or chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell

population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell chemotaxis.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis)consist of assays

10 that measure the ability of a protein to induce the migration of cells across a membrane as well as the ability of a protein to induce the adhesion of one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: Current Protocols in

15 Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W.Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 6.12, Measurement of alpha and beta Chemokines 6.12.1-6.12.28; Taub et al. J. Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Muller et al Eur. J. Immunol. 25: 1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol. 153: 1762-1768, 1994.

Hemostatic and Thrombolytic Activity

A protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulation disorders (includinghereditary disorders, such as hemophilias) or to enhance coagulation and other hemostatic events in treating wounds resulting from trauma, surgery or other causes. A

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protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as, for example, infarction of cardiac and central nervous system vessels (e.g., stroke).

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The activity of a protein of the invention may, among other means, be measured by the following methods:

Assay for hemostatic and thrombolytic activity include, without limitation, those described in: Linet et al., J. Clin.

10 Pharmacol. 26:131-140, 1986; Burdick et al., Thrombosis Res.
45:413-419, 1987; Humphrey et al., Fibrinolysis 5:71-79 (1991);
Schaub, Prostaglandins 35:467-474, 1988.

Receptor/Ligand Activity

A protein of the present invention may also demonstrate 15 activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands, receptors 20 involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and their ligands) and receptor/ligand pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune responses). 25 Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction. A protein of the present invention (including, without limitation, fragments receptors and ligands) may themselves be useful as inhibitors

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of receptor/ligand interactions.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Suitable assays for receptor-ligand activity include

without limitation those described in:Current Protocols in
Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies,
E.M. Shevach, W.Strober, Pub. Greene Publishing Associates and
Wiley-Interscience (Chapter 7.28, Measurement of Cellular
Adhesion under static conditions 7.28.1-7.28.22), Takai et al.,

Proc. Natl. Acad. Sci. USA 84:6864-6868, 1987; Bierer et al.,
J. Exp. Med. 168:1145-1156, 1988; Rosenstein et al., J. Exp.
Med. 169:149-160 1989; Stoltenborg et al., J. Immunol.
Methods 175:59-68, 1994; Stitt et al., Cell 80:661-670, 1995.

Anti-Inflammatory Activity

Proteins of the present invention may also exhibit 15 anti-inflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example, cell adhesion), by 20 inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can 25 be used to treat inflammatory conditions including chronic or acute conditions), including without limitation inflammation associated with infection (such as septic shock, sepsis or syndrome inflammatory response systemic ischemia-reperfusion injury, endotoxin lethality, arthritis,

complement-mediated hyperacute rejection, nephritis, cytokine or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of ytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material.

Tumor Inhibition Activity

In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth

20 Other Activities

A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in

bone form or shape); effecting biorhythms or caricadic cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or elimination of dietary fat, lipid, 5 protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent 10 behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related 15 diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another 20 material or entity which is cross-reactive with such protein.

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Sequence Table

	(2)	INF	ORMA'	rion	FOR	SEQ	ID 1	10:	1:							
5		(:	i) S	EQUE	NCE (CHAR	ACTE	RIST	ics:							
				(A)	LEN	GTH:	382									
				(B)	TYP	E: Ar	nino	aci	i							
				(D)	TOP	DLOG:	Y: L:	inea	r							
		(.	ii) :	SEQUI	ENCE	KIN	D: P:	rote	in							
10		(.	iii)	HYP	THE:	TICA	L: No	5								
		(vi) (ORIG	INAL	sou	RCE:	•								
				(A)	ORG	ANIS	M: H	ото	sapi	ens						
				(B)	CEL	L KI	ND: I	Live	r							
15				(D)	CLO	NE NA	AME:	HPO:	1263							
		(:	xi) :	SEQUI	ENCE	DES	CRIP!	rion	: SE	Q ID	NO:	1:				
	Met	Gly	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Cys	Ile	Leu	Val	Leu	Cys	Суя
20	1				5					10					15	
	Gly	Ala	Met	Ser	Pro	Pro	Gln	Leu		Leu	Asn	Pro	Ser		Leu	Leu
				20					25					30		
	Ser	Arg		Cys	Asn	Asp	Ser		Val	Leu	Ala	Val		Gly	Phe	Ala
			35					40					45			
25	Leu	Arg	Asp	Ile	Asn	Lys	Asp	Arg	Lys	Asp	Gly	Tyr	Val	Leu	Arg	Let
		50					55					60				
	Asn	Arg	Val	Asn	Asp	Ala	Gln	Glu	Tyr	Arg	Arg	Gly	Gly	Leu	G1y	
	65					70					75					80
	Leu	Phe	Tyr	Leu	Thr	Leu	Asp	Val	Leu	Glu	Thr	Asp	Cys	His	Val	Leu
30					85					90					95	
	Arg	Lys	Lys	Ala	Trp	Gln	Asp	Cys	Gly	Met	Arg	Ile	Phe	Phe	Glu	Ser
				100					105					110		
	Val	Tyr	Gly	Gln	Cys	Lys	Ala	Ile	Phe	Tyr	Met	Asn	Asn	Pro	Ser	Arg
			115					120					125			
35	Val	Leu	Tyr	Leu	Ala	Ala	Tyr	Asn	Cys	Thr	Leu	Arg	Pro	Val	Ser	Lys

Lys Lys Ile Tyr Met Thr Cys Pro Asp Cys Pro Ser Ser Ile Pro Thr

84

	Asp	Ser	Ser	Asn	His	Gln	Val	Leu	Glu	Ala	Ala	Thr	Glu	Ser	Leu	Ala
					165					170					175	
	Lys	Tyr	Asn	Asn	Glu	Asn	Thr	Ser	Lys	Gln	Tyr	Ser	Leu	Phe	Lys	Val
				180					185					190		
5	Thr	Arg	Ala	Ser	Ser	Gln	Trp	Val	Val	Gly	Pro	Ser	Tyr	Phe	Val	Glu
			195					200					205			
	Tyr	Leu	Ile	Lys	Glu	Ser	Pro	Cys	Thr	Lys	Ser	Gln	Ala	Ser	Ser	Cys
		210					215					220				
	Ser	Leu	Gln	Ser	Ser	Asp	Ser	Val	Pro	Val	Gly	Leu	Cys	Lys	Gly	Ser
10	225					230					235					240
	Leu	Thr	Arg	Thr	His	Trp	Glu	Lys	Phe	Val	Ser	Val	Thr	Cys	Asp	Phe
					245					250					255	
	Phe	Glu	Ser	Gln	Ala	Pro	Ala	Thr	Gly	Ser	Glu	Asn	Ser	Ala	Val	Asn
				260					265					270		
15	Gln	Lys	Pro	Thr	Asn	Leu	Pro	Lys	Val	Glu	Glu	Ser	Gln	Gln	Lys	Asn
			275					280					285			
	Thr	Pro	Pro	Thr	Asp	Ser	Pro	Ser	Lys	Ala	Gly	Pro	Arg	Gly	Ser	Val
		290					295					300				
	Gln	Tyr	Leu	Pro	Asp	Leu	Asp	Asp	Lys	Asn	Ser	Gln	Glu	Lys	Gly	
20	305					310					315					320
	Gln	Glu	Ala	Phe	Pro	Val	His	Leu	Asp	Leu	Thr	Thr	Asn	Pro	Gln	Gly
					325					330					335	
	G1u	Thr	Leu	Asp	Ile	Ser	Phe	Leu	Phe	Leu	Glu	Pro	Met		Glu	Lys
				340					345					350		
25	Leu	Val	Val	Leu	Pro	Phe	Pro	Lys	Glu	Lys	Ala	Arg	Thr	Ala	Glu	Cys
			355					360					365			
	Pro	Gly	Pro	Ala	Gln	Asn	Ala	Ser	Pro	Leu	Val	Leu	Pro	Pro		
		370					375					380				

- (2) INFORMATION FOR SEQ ID NO: 2:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 317
 - (B) TYPE: Amino acid
- 35 (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: Protein
 - (iii) HYPOTHETICAL: No

85

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(B) CELL KIND: Liver

(D) CLONE NAME: HP01299

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

	Met	Trp	Leu	Tyr	Leu	Ala	Ala	Phe	Val	Gly	Leu	Tyr	Tyr	Leu	Leu	His
	1				5					10					15	
10	Trp	Tyr	Arg	Glu	Arg	Gln	Val	Val	Ser	His	Leu	Gln	Asp	Lys	Tyr	Val
				20					25					30		
	Phe	Ile	Thr	Gly	Cys	Asp	Ser	Gly	Phe	Gly	Asn	Leu	Leu	Ala	Arg	Gln
			35					40					45			
	Leu	Asp	Ala	Arg	Gly	Leu	Arg	Val	Leu	Ala	Ala	Cys	Leu	Thr	Glu	Lys
15		50					55					60				
	G1y	Ala	Glu	Gln	Leu	Arg	Gly	Gln	Thr	Ser	Asp	Arg	Leu	Glu	Thr	Val
	65					70					75					80
	Thr	Leu	Asp	Val	Thr	Lys	Met	Glu	Ser	Ile	Ala	Ala	Ala	Thr	Gln	Trp
					85					90					95	
20	Val	Lys	Glu	His	Val	Gly	Asp	Arg	Gly	Leu	Trp	G1y	Leu	Val	Asn	Asn
				100					105	,				110		
	Ala	Gly	Ile	Leu	Thr	Pro	Ile	Thr	Leu	Cys	Glu	Trp	Leu	Asn	Thr	Glu
			115					120					125			
	Asp	Ser	Met	Asn	Met	Leu	Lys	Val	Asn	Leu	Ile	Gly	Val	Ile	Gln	Val
25		130					135					140				
	Thr	Leu	Ser	Met	Leu	Pro	Leu	Val	Arg	Arg	Ala	Arg	Gly	Arg	Ile	
	145					150					155					160
	Asn	Val	Ser	Ser	Ile	Leu	Gly	Arg	Val	Ala	Phe	Phe	Val	Gly		Tyr
					165					170					175	
30	Cys	Val	Ser	-	Tyr	Gly	Val	Glu		Phe	Ser	Asp	Ile		Arg	Arg
				180					185					190		_
	Glu	Ile		His	Phe	Gly	Val		Ile	Ser	Ile	Val		Pro	Gly	Tyr
			195					200					205			_
	Phe		Thr	Gly	Met	Thr	Asn	Met	Thr	Gln	Ser		Glu	Arg	Met	Lys
35		210					215					220				
		Ser	Trp	Lys	Glu		Pro	Lys	His	Ile		Glu	Thr	Tyr	Gly	
	225					230					235			_		240
	Gln	Tyr	Phe	Asp	Ala	Leu	Tyr	Asn	Ile	Met	Lys	Glu	Gly	Leu	Leu	Asn

86

250 245 Cys Ser Thr Asn Leu Asn Leu Val Thr Asp Cys Met Glu His Ala Leu 265 Thr Ser Val His Pro Arg Thr Arg Tyr Ser Ala Gly Trp Asp Ala Lys 280 275 Phe Phe Phe Ile Pro Leu Ser Tyr Leu Pro Thr Ser Leu Ala Asp Tyr 295 Ile Leu Thr Arg Ser Trp Pro Lys Pro Ala Gln Ala Val 305 310 315 10 (2) INFORMATION FOR SEQ ID NO: 3: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 296 (B) TYPE: Amino acid 15 (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (B) CELL KIND: Liver (D) CLONE NAME: HP01347 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3: Met Ser Asp Ser Lys Glu Pro Arg Val Gln Gln Leu Gly Leu Leu Gly 15 1 Cys Leu Gly His Gly Ala Leu Val Leu Gln Leu Leu Ser Phe Met Leu 30 25 Leu Ala Gly Val Leu Val Ala Ile Leu Val Gln Val Ser Lys Val Pro 45 40 Ser Ser Leu Ser Gln Glu Gln Ser Glu Gln Asp Ala Ile Tyr Gln Asn 55 35 Leu Thr Gln Leu Lys Ala Ala Val Gly Glu Leu Ser Glu Lys Ser Lys 75 70 65 Leu Gln Glu Ile Tyr Gln Glu Leu Thr Gln Leu Lys Ala Ala Val Gly

90

85

	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr
				100					105					110		
	Arg	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln
			115					120					125			
5	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu
		130					135					140				
	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu
	145					150					155					160
	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile
10					165					170					175	
	Tyr	Gln	Glu	Leu	Thr	Glu	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu
				180					185					190		
	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Gln	Leu	Lys	Ala
			195					200					205			
15	Ala	Val	Gly	Glu	Leu	Pro	Asp	Gln	Ser	Lys	Gln	Gln	Gln	Ile	Tyr	Gln
		210					215					220				
	Glu	Leu	Thr	Asp	Leu	Lys	Thr	Ala	Phe	Glu	Arg	Leu	Cys	Arg	His	Cys
	225					230					235					240
	Pro	Lys	Asp	Trp	Thr	Phe	Phe	Gln	Gly	Asn	Cys	Tyr	Phe	Met		Asn
20					245			٠		250					255	
	Ser	Gln	Arg	Asn	Trp	His	Asp	Ser	Val	Thr	Ala	Cys	Gln	G1u	Val	Arg
				260					265					270		
	Ala	Gln		Val	Val	Ile	Lys		Ala	Glu	Glu	Gln		Pro	Ala	Val
			275					280					285			
25	Leu		Gln	Trp	Arg	Thr		Gln								
		290					295									

- (2) INFORMATION FOR SEQ ID NO: 4:
- 30 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 197
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: Protein
- 35 (iii) HYPOTHETICAL: No
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens

- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP01440

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

5

Met Cys Thr Gly Lys Cys Ala Arg Cys Val Gly Leu Ser Leu Ile Thr 10 Leu Cys Leu Val Cys Ile Val Ala Asn Ala Leu Leu Leu Val Pro Asn 25 10 Gly Glu Thr Ser Trp Thr Asn Thr Asn His Leu Ser Leu Gln Val Trp 40 Leu Met Gly Gly Phe Ile Gly Gly Leu Met Val Leu Cys Pro Gly Ile Ala Ala Val Arg Ala Gly Gly Lys Gly Cys Cys Gly Ala Gly Cys 75 70 15 Cys Gly Asn Arg Cys Arg Met Leu Arg Ser Val Phe Ser Ser Ala Phe 85 90 Gly Val Leu Gly Ala Ile Tyr Cys Leu Ser Val Ser Gly Ala Gly Leu 105 20 Arg Asn Gly Pro Arg Cys Leu Met Asn Gly Glu Trp Gly Tyr His Phe 120 115 Glu Asp Thr Ala Gly Ala Tyr Leu Leu Asn Arg Thr Leu Trp Asp Arg 135 Cys Glu Ala Pro Pro Arg Val Val Pro Trp Asn Val Thr Leu Phe Ser 25 145 150 Leu Leu Val Ala Ala Ser Cys Leu Glu Ile Val Leu Cys Gly Ile Gln 170 Leu Val Asn Ala Thr Ile Gly Val Phe Cys Gly Asp Cys Arg Lys Lys 190 180 185

30 Gln Asp Thr Pro His

195

- (2) INFORMATION FOR SEQ ID NO: 5:
 - (i) SEQUENCE CHARACTERISTICS:

35 (A) LENGTH: 221

- (B) TYPE: Amino acid
- (D) TOPOLOGY: Linear
- (ii) SEQUENCE KIND: Protein

(iii) HYPOTHETICAL: No

(vi) ORIGINAL SOURCE:

5

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
 - (D) CLONE NAME: HP01526

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

10	Met	Glu	Ala	G1y	Gly	Phe	Leu	Asp	Ser	Leu	Ile	Tyr	Gly	Ala	Cys	Val
	1				5					10					15	
	Val	Phe	Thr	Leu	Gly	Met	Phe	Ser	Ala	Gly	Leu	Ser	Asp	Leu	Arg	His
				20					25					30		
	Met	Arg	Met	Thr	Arg	Ser	Val	Asp	Asn	Val	Gln	Phe	Leu	Pro	Phe	Leu
15			35					40					45			
	Thr	Thr	Glu	Val	Asn	Asn	Leu	Gly	Trp	Leu	Ser	Tyr	Gly	Ala	Leu	Lys
		50					55					60				
	Gly	Asp	Gly	Ile	Leu	Ile	Val	Val	Asn	Thr	Val	Gly	Ala	Ala	Leu	Gln
	65					70					75					80
20	Thr	Leu	Tyr	Ile	Leu	Ala	Tyr	Leu	His	Tyr	Cys	Pro	Arg	Lys	Arg	Val
-					85					90					95	
	Val	Leu	Leu	Gln	Thr	Ala	Thr	Leu	Leu	Gly	Val	Leu	Leu	Leu	Gly	Tyr
				100					105					110		
	Gly	Tyr	Phe	Trp	Leu	Leu	Val	Pro	Asn	Pro	Glu	Ala	Arg	Leu	Gln	Gln
25			115					120					125			
	Leu	Gly	Leu	Phe	Cys	Ser	Val	Phe	Thr	Ile	Ser	Met	Tyr	Leu	Ser	Pro
		130					135					140				
	Leu	Ala	Asp	Leu	Ala	Lys	Val	Ile	Gln	Thr	Lys	Ser	Thr	Gln	Cys	Leu
	145					150					155					160
30	Ser	Tyr	Pro	Leu	Thr	Ile	Ala	Thr	Leu	Leu	Thr	Ser	Ala	Ser	Trp	Cys
					165					170					175	
	Leu	Tyr	Gly	Phe	Arg	Leu	Arg	Asp	Pro	Tyr	Ile	Met	Val	Ser	Asn	Phe
				180					185					190		
	Pro	G1y	Ile	Val	Thr	Ser	Phe	Ile	Arg	Phe	Trp	Leu	Phe	Trp	Lys	Tyr
35			195					200					205			
	Pro	Gln	Glu	Gln	Asp	Arg	Asn	Tyr	Trp	Leu	Leu	Gln	Thr			
		210					215					220				

- 90 (2) INFORMATION FOR SEQ ID NO: 6: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 251 (B) TYPE: Amino acid (D) TOPOLOGY: Linear 5 (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens 10 (B) CELL KIND: Stomach cancer (D) CLONE NAME: HP10230 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6: 15 Met Ser Asp Ile Gly Asp Trp Phe Arg Ser Ile Pro Ala Ile Thr Arg 1 Tyr Trp Phe Ala Ala Thr Val Ala Val Pro Leu Val Gly Lys Leu Gly 25 20 Leu Ile Ser Pro Ala Tyr Leu Phe Leu Trp Pro Glu Ala Phe Leu Tyr 40 35 Arg Phe Gln Ile Trp Arg Pro Ile Thr Ala Thr Phe Tyr Phe Pro Val 60 55 Gly Pro Gly Thr Gly Phe Leu Tyr Leu Val Asn Leu Tyr Phe Leu Tyr 75 25 65 70 Gln Tyr Ser Thr Arg Leu Glu Thr Gly Ala Phe Asp Gly Arg Pro Ala 85 90 Asp Tyr Leu Phe Met Leu Leu Phe Asn Trp Ile Cys Ile Val Ile Thr 105 100 30 Gly Leu Ala Met Asp Met Gln Leu Leu Met Ile Pro Leu Ile Met Ser 120 Val Leu Tyr Val Trp Ala Gln Leu Asn Arg Asp Met Ile Val Ser Phe 135 130 Trp Phe Gly Thr Arg Phe Lys Ala Cys Tyr Leu Pro Trp Val Ile Leu
 - Asn Leu Val Gly His Leu Tyr Phe Phe Leu Met Phe Arg Tyr Pro Met

Gly Phe Asn Tyr Ile Ile Gly Gly Ser Val Ile Asn Glu Leu Ile Gly

150

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91 190 185 180 Asp Leu Gly Gly Arg Asn Phe Leu Ser Thr Pro Gln Phe Leu Tyr Arg 200 Trp Leu Pro Ser Arg Arg Gly Gly Val Ser Gly Phe Gly Val Pro Pro 220 215 210 5 Ala Ser Met Arg Arg Ala Ala Asp Gln Asn Gly Gly Gly Arg His 230 235 225. Asn Trp Gly Gln Gly Phe Arg Leu Gly Asp Gln 250 245 10 (2) INFORMATION FOR SEQ ID NO: 7: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 106 (B) TYPE: Amino acid 15 (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: 20 (A) ORGANISM: Homo sapiens (B) CELL KIND: Epidermoid carcinoma (C) CELL LINE: KB (D) CLONE NAME: HP10389 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7: Met Ala Thr Pro Gly Pro Val Ile Pro Glu Val Pro Phe Glu Pro Ser 1 5 10 30 Lys Pro Pro Val Ile Glu Gly Leu Ser Pro Thr Val Tyr Arg Asn Pro 25 Glu Ser Phe Lys Glu Lys Phe Val Arg Lys Thr Arg Glu Asn Pro Val 35 40 Val Pro Ile Gly Cys Leu Ala Thr Ala Ala Ala Leu Thr Tyr Gly Leu 55 35 Tyr Ser Phe His Arg Gly Asn Ser Gln Arg Ser Gln Leu Met Met Arg 80

70

Thr Arg Ile Ala Ala Gln Gly Phe Thr Val Ala Ala Ile Leu Leu Gly

65

92

85 90 95

Leu Ala Val Thr Ala Met Lys Ser Arg Pro 100 105

5

- (2) INFORMATION FOR SEQ ID NO: 8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 78
- 10 (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: Protein
 - (iii) HYPOTHETICAL: No
 - 15 (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (B) CELL KIND: Stomach cancer
 - (D) CLONE NAME: HP10408
 - 20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

Met Gly Ser Gly Leu Pro Leu Val Leu Leu Leu Thr Leu Leu Gly Ser

5 10 1

Ser His Gly Thr Gly Pro Gly Met Thr Leu Gln Leu Lys Leu Lys Glu

25 20 25 30

Ser Phe Leu Thr Asn Ser Ser Tyr Glu Ser Ser Phe Leu Glu Leu Leu

35 40 45

Glu Lys Leu Cys Leu Leu His Leu Pro Ser Gly Thr Ser Val Thr

50 55 6

30 Leu His His Ala Arg Ser Gln His His Val Val Cys Asn Thr

65 70 75

- (2) INFORMATION FOR SEQ ID NO: 9:
- 35 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 314
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear

- (ii) SEQUENCE KIND: Protein
- (iii) HYPOTHETICAL: No

(vi) ORIGINAL SOURCE:

5 (A) ORGANISM: Homo sapiens

- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10412

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9: 10 Met Val Ala Pro Val Trp Tyr Leu Val Ala Ala Ala Leu Leu Val Gly 1 Phe Ile Leu Phe Leu Thr Arg Ser Arg Gly Arg Ala Ala Ser Ala Gly 15 Gln Glu Pro Leu His Asn Glu Glu Leu Ala Gly Ala Gly Arg Val Ala 40 Gln Pro Gly Pro Leu Glu Pro Glu Glu Pro Arg Ala Gly Gly Arg Pro Arg Arg Arg Asp Leu Gly Ser Arg Leu Gln Ala Gln Arg Arg Ala 75 20 70 Gln Arg Val Ala Trp Ala Glu Ala Asp Glu Asn Glu Glu Glu Ala Val 90 Ile Leu Ala Gln Glu Glu Gly Val Glu Lys Pro Ala Glu Thr His 110 105 25 Leu Ser Gly Lys Ile Gly Ala Lys Lys Leu Arg Lys Leu Glu Glu Lys 120 Gln Ala Arg Lys Ala Gln Arg Glu Ala Glu Glu Ala Glu Arg Glu Glu 135 140 Arg Lys Arg Leu Glu Ser Gln Arg Glu Ala Glu Trp Lys Lys Glu Glu 155 150 30 145 Glu Arg Leu Arg Leu Glu Glu Glu Glu Glu Glu Glu Glu Arg Lys 170 165 Ala Arg Glu Glu Gln Ala Gln Arg Glu His Glu Glu Tyr Leu Lys Leu 185 35 Lys Glu Ala Phe Val Val Glu Glu Glu Gly Val Gly Glu Thr Met Thr 195 200 Glu Glu Gln Ser Gln Ser Phe Leu Thr Glu Phe Ile Asn Tyr Ile Lys 220 210 215

94

Gln Ser Lys Val Val Leu Leu Glu Asp Leu Ala Ser Gln Val Gly Leu 235 230 225 Arg Thr Gln Asp Thr Ile Asn Arg Ile Gln Asp Leu Leu Ala Glu Gly 250 245 5 Thr Ile Thr Gly Val Ile Asp Asp Arg Gly Lys Phe Ile Tyr Ile Thr 265 260 Pro Glu Glu Leu Ala Ala Val Ala Asn Phe Ile Arg Gln Arg Gly Arg 280 Val Ser Ile Ala Glu Leu Ala Gln Ala Ser Asn Ser Leu Ile Ala Trp 300 10 295 290 Gly Arg Glu Ser Pro Ala Gln Ala Pro Ala 305 310

- 15 (2) INFORMATION FOR SEQ ID NO: 10:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 195
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
- 20 (ii) SEQUENCE KIND: Protein
 - (iii) HYPOTHETICAL: No
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- 25 (B) CELL KIND: Stomach cancer
 - (D) CLONE NAME: HP10413
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:

75 70 65 Phe Asp Gly Val Gln Asp Pro Arg Ile Leu Met Ala Ile Asn Gly Lys 85 90 Val Phe Asp Val Thr Lys Gly Arg Lys Phe Tyr Gly Pro Glu Gly Pro 105 5 100 Tyr Gly Val Phe Ala Gly Arg Asp Ala Ser Arg Gly Leu Ala Thr Phe 120 Cys Leu Asp Lys Glu Ala Leu Lys Asp Glu Tyr Asp Asp Leu Ser Asp - 140 135 130 10 Leu Thr Ala Ala Gln Gln Glu Thr Leu Ser Asp Trp Glu Ser Gln Phe 155 150 Thr Phe Lys Tyr His His Val Gly Lys Leu Leu Lys Glu Gly Glu Glu 170 165 Pro Thr Val Tyr Ser Asp Glu Glu Glu Pro Lys Asp Glu Ser Ala Arg 185 180 15 Lys Asn Asp 195 20 (2) INFORMATION FOR SEQ ID NO: 11: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 462 (B) TYPE: Amino acid (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein 25 (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (B) CELL KIND: Stomach cancer 30 (D) CLONE NAME: HP10415 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11: 35 Met Leu Asp Phe Ala Ile Phe Ala Val Thr Phe Leu Leu Ala Leu Val 1 Gly Ala Val Leu Tyr Leu Tyr Pro Ala Ser Arg Gln Ala Ala Gly Ile 25 30 20

	Pro	Gly	Ile	Thr	Pro	Thr	Glu	Glu	Lys	Asp	Gly	Asn	Leu	Pro	Asp	lle
			35					40					45			
	Val	Asn	Ser	Gly	Ser	Leu	His	Glu	Phe	Leu	Val	Asn	Leu	His	Glu	Arg
		50					55					60				
5	Tyr	Gly	Pro	Val	Val	Ser	Phe	Trp	Phe	Gly	Arg	Arg	Leu	Val	Val	Ser
	65					70					75					80
	Leu	Gly	Thr	Val	Asp	Val	Leu	Lys	Gln	His	Ile	Asn	Pro	Asn	Lys	Thr
					85					90					95	
	Leu	Asp	Pro	Phe	Glu	Thr	Met	Leu	Lys	Ser	Leu	Leu	Arg	Tyr	Gln	Ser
10				100					105					110		
	Gly	Gly	Gly	Ser	Val	Ser	Glu	Asn	His	Met	Arg	Lys	Lys	Leu	Tyr	Glu
	•		115					120					125			
	Asn	Gly	Val	Thr	Asp	Ser	Leu	Lys	Ser	Asn	Phe	Ala	Leú	Leu	Leu	Lys
		130					135					140				
15	Leu	Ser	Glu	Glu	Leu	Leu	Asp	Lys	Trp	Leu	Ser	Tyr	Pro	Glu	Thr	Gln
	145					150					155					160
	His	Val	Pro	Leu	Ser	Gln	His	Met	Leu	Gly	Phe	Ala	Met	Lys	Ser	Val
					165					170					175	
	Thr	Gln	Met	Val	Met	Gly	Ser	Thr	Phe	Glu	Asp	Asp	Gln	Glu	Val	Ile
20				180					185					190		
	Arg	Phe	Gln	Lys	Asn	His	Gly	Thr	Val	Trp	Ser	Glu	Ile	Gly	Lys	Gly
			195					200					205			
	Phe	Leu	Asp	Gly	Ser	Leu	Asp	Lys	Asn	Met	Thr	Arg	Lys	Lys	Gln	Tyr
		210					215					220				
25	Glu	Asp	Ala	Leu	Met	Gln	Leu	Glu	Ser	Val	Leu	Arg	Asn	Ile	Ile	Lys
	225					230					235					240
	Glu	Arg	Lys	Gly	Arg	Asn	Phe	Ser	Gln	His	Ile	Phe	Ile	Asp	Ser	Leu
					245					250					255	
	Val	Gln	Gly	Asn	Leu	Asn	Asp	Gln	Gln	Ile	Leu	Glu	Asp	Ser	Met	Ile
30				260					265					270		
	Phe	Ser	Leu	Ala	Ser	Cys	Ile	Ile	Thr	Ala	Lys	Leu	Cys	Thr	Trp	Ala
			275					280					285			
	Ile	Cys	Phe	Leu	Thr	Thr	Ser	G1u	Glu	Val	Gln	Lys	Lys	Leu	Tyr	Glu
		290					295					300				
35	Glu	Ile	Asn	Gln	Val	Phe	Gly	Asn	Gly	Pro	Val	Thr	Pro	Glu	Lys	Ile
	305					310					315					320
	Glu	Gln	Leu	Arg	Tyr	Cys	Gln	His	Val	Leu	Cys	Glu	Thr	Val	Arg	Thr
					325					330					335	

Ala Lys Leu Thr Pro Val Ser Ala Gln Leu Gln Asp Ile Glu Gly Lys 345 Ile Asp Arg Phe Ile Ile Pro Arg Glu Thr Leu Val Leu Tyr Ala Leu 360 5 Gly Val Val Leu Gln Asp Pro Asn Thr Trp Pro Ser Pro His Lys Phe 380 375 370 Asp Pro Asp Arg Phe Asp Asp Glu Leu Val Met Lys Thr Phe Ser Ser 395 390 Leu Gly Phe Ser Gly Thr Gln Glu Cys Pro Glu Leu Arg Phe Ala Tyr 405 410 10 Met Val Thr Thr Val Leu Leu Ser Val Leu Val Lys Arg Leu His Leu 430 425 Leu Ser Val Glu Gly Gln Val Ile Glu Thr Lys Tyr Glu Leu Val Thr 440 15 Ser Ser Arg Glu Glu Ala Trp Ile Thr Val Ser Lys Arg Tyr 460 455

- (2) INFORMATION FOR SEQ ID NO: 12:
- 20 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 247
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: Protein
- 25 (iii) HYPOTHETICAL: No
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (B) CELL KIND: Stomach cancer
- 30 (D) CLONE NAME: HP10419
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

													45			
			35					40							_	
	Ala	Ser	Val	Val	Trp	Phe		Leu	Val	His	Val	Thr	Asp	Arg	Ser	Asp
		50					55					60				
	Ala	Arg	Leu	Gln	Tyr	Gly	Leu	Leu	Ile	Phe	Gly	Ala	Ala	Val	Ser	Val
5	65					70					75					80
	Leu	Leu	Gln	Glu	Val	Phe	Arg	Phe	Ala	Tyr	Tyr	Lys	Leu	Leu	Lys	Lys
					85					90					95	
	Ala	Asp	Glu	Gly	Leu	Ala	Ser	Leu	Ser	Glu	Asp	Gly	Arg	Ser	Pro	Ile
				100					105					110		
10	Ser	Ile	Arg	Gln	Met	Ala	Tyr	Val	Ser	Gly	Leu	Ser	Phe	Gly	Ile	Ile
			115					120					125			
	Ser	Gly	Val	Phe	Ser	Va1	Ile	Asn	Ile	Leu	Ala	Asp	Ala	Leu	Gly	Pro
		130					135					140				
	Gly	Val	Val	Gly	Ile	His	Gly	Asp	Ser	Pro	Tyr	Tyr	Phe	Leu	Thr	Ser
15	145					150					155					160
	Ala	Phe	Leu	Thr	Ala	Ala	Ile	Ile	Leu	Leu	His	Thr	Phe	Trp	Gly	Val
					165					170					175	
	Val	Phe	Phe	Asp	Ala	Cys	Glu	Arg	Arg	Arg	Tyr	Trp	Ala	Leu	Gly	Leu
				180		-			185					190		
20	Val	Val	Gly	Ser	His	Leu	Leu	Thr	Ser	Gly	Leu	Thr	Phe	Leu	Asn	Pro
			195					200					205			
	Trp	Tyr	Glu	Ala	Ser	Leu	Leu	Pro	Ile	Tyr	Ala	Val	Thr	Val	Ser	Met
	•	210					215			-		220				
	G1 v		Tro	Ala	Phe	Ile	Thr	Ala	Glv	Glv	Ser	Leu	Arg	Ser	Ile	Gln
25	225					230					235		Ŭ			240
2.5		Sar	T 011	Leu	Cvc		Asn									
	Ar 8	Der	Dea	Dea	245	ພງວ	p									
					243											

- 30 (2) INFORMATION FOR SEQ ID NO: 13:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 113
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
- 35 (ii) SEQUENCE KIND: Protein
 - (iii) HYPOTHETICAL: No
 - (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10424
- 5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

Met Asn Phe Tyr Leu Leu Leu Ala Ser Ser Ile Leu Cys Ala Leu Ile 1 5 10 15

Val Phe Trp Lys Tyr Arg Arg Phe Gln Arg Asn Thr Gly Glu Met Ser

10 20 25 30

Ser Asn Ser Thr Ala Leu Ala Leu Val Arg Pro Ser Ser Ser Gly Leu
35 40 45

Ile Asn Ser Asn Thr Asp Asn Asn Leu Ala Val Tyr Asp Leu Ser Arg
50 55 60

15 Asp Ile Leu Asn Asn Phe Pro His Ser Ile Ala Arg Gln Lys Arg Ile 65 70 75 80

Leu Val Asn Leu Ser Met Val Glu Asn Lys Leu Val Glu Leu Glu His
85 90 95

Thr Leu Leu Ser Lys Gly Phe Arg Gly Ala Ser Pro His Arg Lys Ser 20 100 105 110

Thr

- (2) INFORMATION FOR SEQ ID NO: 14:
- 25 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 365
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: Protein
- 30 (iii) HYPOTHETICAL: No
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (B) CELL KIND: Epidermoid carcinoma
- 35 (C) CELL LINE: KB
 - (D) CLONE NAME: HP10428
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

	Met	Gly	Arg	Trp	Ala	Leu	Asp	Val	Ala	Phe	Leu	Trp	Lys	Ala	Val	Leu
	1				5					10					15	
	Thr	Leu	Gly	Leu	Val	Leu	Leu	Tyr	Tyr	Cys	Phe	Ser	Ile	Gly	Ile	Thr
				20					25					30		
5	Phe	Tyr	Asn	Lys	Trp	Leu	Thr	Lys	Ser	Phe	His	Phe	Pro	Leu	Phe	Met
			35					40					45			
	Thr	Met	Leu	His	Leu	Ala	Val	Ile	Phe	Leu	Phe	Ser	Ala	Leu	Ser	Arg
		50					55					60				
	Ala	Leu	Val	Gln	Cys	Ser	Ser	His	Arg	Ala	Arg	Val	Val	Leu	Ser	Trp
10	65					70					75					80
	Ala	Asp	Tyr	Leu	Arg	Arg	Val	Ala	Pro	Thr	Ala	Leu	Ala	Thr	Ala	Leu
					85					90					95	
	Asp	Val	G1y	Leu	Ser	Asn	Trp	Ser	Phe	Leu	Tyr	Val	Thr	Val	Ser	Leu
				100					105					110		
15	Tyr	Thr	Met	Thr	Lys	Ser	Ser	Ala	Val	Leu	Phe	Ile	Leu	Ile	Phe	Ser
			115					120					125			
	Leu	Ile	Phe	Lys	Leu	Glu	Glu	Leu	Arg	Ala	Ala	Leu	Val	Leu	Val	Val
		130					135					140				
	Leu	Leu	Ile	Ala	Gly	Gly	Leu	Phe	Met	Phe	Thr	Tyr	Lys	Ser	Thr	Gln
20	145					150					155					160
	Phe	Asn	Val	Glu	Gly	Phe	Ala	Leu	Val	Leu	Gly	Ala	Ser	Phe	Ile	Gly
					165					170					175	
	Gly	Ile	Arg	Trp	Thr	Leu	Thr	Gln	Met	Leu	Leu	Gln	Lys	Ala	Glu	Lev
				180					185					190		
25	Gly	Leu	Gln	Asn	Pro	Ile	Asp	Thr	Met	Phe	His	Leu	Gln	Pro	Leu	Met
			195					200					205			
	Phe	Leu	Gly	Leu	Phe	Pro	Leu	Phe	Ala	Val	Phe	Glu	Gly	Leu	His	Lev
		210					215					220				
	Ser	Thr	Ser	Glu	Lys	Ile	Phe	Arg	Phe	Gln	Asp	Thr	Gly	Leu	Leu	
30	225					230					235					240
	Arg	Va1	Leu	Gly	Ser	Ļeu	Phe	Leu	Gly	Gly	Ile	Leu	Ala	Phe	Gly	Let
					245					250					255	
	Gly	Phe	Ser	Glu	Phe	Leu	Leu	Val	Ser	Arg	Thr	Ser	Ser	Leu	Thr	Lei
				260					265					270		
35	Ser	Ile	Ala	Gly	Ile	Phe	Lys	Glu	Val	Cys	Thr	Leu	Leu	Leu	Ala	Ala
			275					280					285			
	His	Leu	Leu	Gly	Asp	Gln	Ile	Ser	Leu	Leu	Asn	Trp	Leu	Gly	Phe	Ala
		200					295					300				

101

Leu Cys Leu Ser Gly Ile Ser Leu His Val Ala Leu Lys Ala Leu His 310 315 305 Ser Arg Gly Asp Gly Gly Pro Lys Ala Leu Lys Gly Leu Gly Ser Ser 325 330 5 Pro Asp Leu Glu Leu Leu Arg Ser Ser Gln Arg Glu Glu Gly Asp 350 340 345 Asn Glu Glu Glu Tyr Phe Val Ala Gln Gly Gln Gln 360 365 10 (2) INFORMATION FOR SEQ ID NO: 15: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 226 (B) TYPE: Amino acid 15 (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: 20 (A) ORGANISM: Homo sapiens (B) CELL KIND: Stomach cancer (D) CLONE NAME: HP10429 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15: 25 Met Pro Thr Thr Lys Lys Thr Leu Met Phe Leu Ser Ser Phe Phe Thr 10 Ser Leu Gly Ser Phe Ile Val Ile Cys Ser Ile Leu Gly Thr Gln Ala 20 25 Trp Ile Thr Ser Thr Ile Ala Val Arg Asp Ser Ala Ser Asn Gly Ser 40 Ile Phe Ile Thr Tyr Gly Leu Phe Arg Gly Glu Ser Ser Glu Glu Leu 50 55 Ser His Gly Leu Ala Glu Pro Lys Lys Phe Ala Val Leu Glu Ile 35 70 75 Leu Asn Asn Ser Ser Gln Lys Thr Leu His Ser Val Thr Ile Leu Phe 90 85 Leu Val Leu Ser Leu Ile Thr Ser Leu Leu Ser Ser Gly Phe Thr Phe

102

110 105 100 Tyr Asn Ser Ile Ser Asn Pro Tyr Gln Thr Phe Leu Gly Pro Thr Gly 120 Val Tyr Thr Trp Asn Gly Leu Gly Ala Ser Phe Val Phe Val Thr Met 140 135 5 130 Ile Leu Phe Val Ala Asn Thr Gln Ser Asn Gln Leu Ser Glu Glu Leu 150 155 Phe Gln Met Leu Tyr Pro Ala Thr Thr Ser Lys Gly Thr Thr His Ser 170 10 Tyr Gly Tyr Ser Phe Trp Leu Ile Leu Leu Val Ile Leu Leu Asn Ile 185 Val Thr Val Thr Ile Ile Ile Phe Tyr Gln Lys Ala Arg Tyr Gln Arg 205 195 200 Lys Gln Glu Gln Arg Lys Pro Met Glu Tyr Ala Pro Arg Asp Gly Ile 215 220 210 15 Leu Phe 225

- 20 (2) INFORMATION FOR SEQ ID NO: 16:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 129
 - (B) TYPE: Amino acid
 - (D) TOPOLOGY: Linear
- 25 (ii) SEQUENCE KIND: Protein
 - (iii) HYPOTHETICAL: No
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- 30 (B) CELL KIND: Liver
 - (D) CLONE NAME: HP10432
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:
- 35 Met Ala Arg Gly Ser Leu Arg Arg Leu Leu Arg Leu Leu Val Leu Gly

 1 5 10 15

 Leu Trp Leu Ala Leu Leu Arg Ser Val Ala Gly Glu Gln Ala Pro Gly

 20 25 30

103

Thr Ala Pro Cys Ser Arg Gly Ser Ser Trp Ser Ala Asp Leu Asp Lys 40 35 Cys Met Asp Cys Ala Ser Cys Arg Ala Arg Pro His Ser Asp Phe Cys 55 5 Leu Gly Cys Ala Ala Ala Pro Pro Ala Pro Phe Arg Leu Leu Trp Pro 70 75 65 Ile Leu Gly Gly Ala Leu Ser Leu Thr Phe Val Leu Gly Leu Leu Ser 85 Gly Phe Leu Val Trp Arg Arg Cys Arg Arg Arg Glu Lys Phe Thr Thr 105 10 100 Pro Ile Glu Glu Thr Gly Gly Glu Gly Cys Pro Ala Val Ala Leu Ile 120 125 115 Gln 15 (2) INFORMATION FOR SEQ ID NO: 17: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 163 20 (B) TYPE: Amino acid (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: -25 (A) ORGANISM: Homo sapiens (B) CELL KIND: Liver (D) CLONE NAME: HP10433 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17: Met Arg Arg Leu Leu Ile Pro Leu Ala Leu Trp Leu Gly Ala Val Gly 15 10 1 5 Val Gly Val Ala Glu Leu Thr Glu Ala Gln Arg Arg Gly Leu Gln Val 35 25 Ala Leu Glu Glu Phe His Lys His Pro Pro Val Gln Trp Ala Phe Gln 45 40

Glu Thr Ser Val Glu Ser Ala Val Asp Thr Pro Phe Pro Ala Gly Ile

104

60 50 55 Phe Val Arg Leu Glu Phe Lys Leu Gln Gln Thr Ser Cys Arg Lys Arg 75 70 Asp Trp Lys Lys Pro Glu Cys Lys Val Arg Pro Asn Gly Arg Lys Arg 5 90 Lys Cys Leu Ala Cys Ile Lys Leu Gly Ser Glu Asp Lys Val Leu Gly 105 Arg Leu Val His Cys Pro Ile Glu Thr Gln Val Leu Arg Glu Ala Glu 125 115 120 10 Glu His Gln Glu Thr Gln Cys Leu Arg Val Gln Arg Ala Gly Glu Asp 135 Pro His Ser Phe Tyr Phe Pro Gly Gln Phe Ala Phe Ser Lys Ala Leu 155 145 150 Pro Arg Ser 15 (2) INFORMATION FOR SEQ ID NO: 18: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 193 20 (B) TYPE: Amino acid (D) TOPOLOGY: Linear (ii) SEQUENCE KIND: Protein (iii) HYPOTHETICAL: No (vi) ORIGINAL SOURCE: 25 (A) ORGANISM: Homo sapiens (B) CELL KIND: Stomach cancer (D) CLONE NAME: HP10480 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18: 30 Met Ile Arg Cys Gly Leu Ala Cys Glu Arg Cys Arg Trp Ile Leu Pro 5 10 1 Leu Leu Leu Ser Ala Ile Ala Phe Asp Ile Ile Ala Leu Ala Gly 25 35 Arg Gly Trp Leu Gln Ser Ser Asp His Gly Gln Thr Ser Ser Leu Trp 35 40 Trp Lys Cys Ser Gln Glu Gly Gly Gly Ser Gly Ser Tyr Glu Glu Gly

	Cys	G1n	Ser	Leu	Met	Glu	Tyr	Ala	Trp	Gly	Arg	Ala	Ala	Ala	Ala	Met		
	65					70					75					80		
	Leu	Phe	Cys	Gly	Phe	Ile	Ile	Leu	Val	Ile	Cys	Phe	Ile	Leu	Ser	Phe		
5					85					90					95			
	Phe	Ala	Leu	Cys	Gly	Pro	Gln	Met	Leu	Val	Phe	Leu	Arg	Val	Ile	Gly		
				100					105					110				
	Gly	Leu	Leu	Ala	Leu	Ala	Ala	Val	Phe	Gln	Ile	Ile	Ser	Leu	Val	Ile		
			115					120					125					
L O	Tyr	Pro	Val	Lys	Tyr	Thr	Gln	Thr	Phe	Thr	Leu	His	Ala	Asn	Arg	Ala		
		130					135					140						
	Val	Thr	Tyr	Ile	Tyr	Asn	Trp	Ala	Tyr	Gly	Phe	Gly	Trp	Ala	Ala	Thr		
	145					150					155					160		
	Ile	Ile	Leu	Ile	Gly	Cys	Ala	Phe	Phe	Phe	Cys	Cys	Leu	Pro	Asn	Tyr		
15					165					170					175			
	Glu	Asp	Asp	Leu	Leu	Gly	Asn	Ala	Lys	Pro	Arg	Tyr	Phe	Tyr	Thr	Ser		
	•			180					185					190				
	Ala																	
	•																	
20																		
	(2)					-		NO: 1										
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		(.	T T J	sEQU.	ENCE	KIN.	υ; c.	DNA 1	го ши	KNA								
		(wil .	ORIG	TNAT	SOIT	· ਜ) 9											
30		`	VI)					ото	sani	ens								
50								Line		C113								
				, ,				HPO:										
				(2)	020	.,												
		(xi)	SEOU	ENCE	DES	CRTP	TION	: SE	O ID	NO:	19:						
35		``	,	DDQO	2.,02	000				ų								
	ATG	GTC	TGC	TCCT	TCCC	CT G	GCAC'	TCTG	С АТ	CCTA	GTCC	TGT	GCTG	CGG	AGCA	ATGTCT		60
																TCCGAT	1	.20
																GGCTAT		.80
						•			•									
										*		-		-				

	GTGCTGAGAC	TCAACCGAGT	GAACGACGCC	CAGGAATACA	GACGGGGTGG	CCTGGGATCT	240
	CTGTTCTATC	TTACACTGGA	TGTGCTAGAG	ACTGACTGCC	ATGTGCTCAG	AAAGAAGGCA	300
	TGGCAAGACT	GTGGAATGAG	GATATTTTT	GAATCAGTTT	ATGGTCAATG	CAAAGCAATA	360
	TTTTATATGA	ACAACCCAAG	TAGAGTTCTC	TATTTAGCTG	CTTATAACTG	TACTCTTCGC	420
5	CCAGTTTCAA	AAAAAAAGAT	TTACATGACG	TGCCCTGACT	GCCCAAGCTC	CATACCCACT	480
	GACTCTTCCA	ATCACCAAGT	GCTGGAGGCT	GCCACCGAGT	CTCTTGCGAA	ATACAACAAT	540
	GAGAACACAT	CCAAGCAGTA	TTCTCTCTTC	AAAGTCACCA	GGGCTTCTAG	CCAGTGGGTG	600
	GTCGGCCCTT	CTTACTTTGT	GGAATACTTA	ATTAAAGAAT	CACCATGTAC	TAAATCCCAG	660
	GCCAGCAGCT	GTTCACTTCA	GTCCTCCGAC	TCTGTGCCTG	TTGGTCTTTG	CAAAGGTTCT	720
LO	CTGACTCGAA	CACACTGGGA	AAAGTTTGTC	TCTGTGACTT	GTGACTTCTT	TGAATCACAG	780
	GCTCCAGCCA	CTGGAAGTGA	AAACTCTGCT	GTTAACCAGA	AACCTACAAA	CCTTCCCAAG	840
	GTGGAAGAAT	CCCAGCAGAA	AAACACCCCC	CCAACAGACT	CCCCTCCAA	AGCTGGGCCA	900
	AGAGGATCTG	TCCAATATCT	TCCTGACTTG	GATGATAAAA	ATTCCCAGGA	AAAGGGCCCT	960
	CAGGAGGCCT	TTCCTGTGCA	TCTGGACCTA	ACCACGAATC	CCCAGGGAGA	AACCCTGGAT	1020
15	ATTTCCTTCC	TCTTCCTGGA	GCCTATGGAG	GAGAAGCTGG	TTGTCCTGCC	TTTCCCCAAA	1080
	GAAAAAGCAC	GCACTGCTGA	GTGCCCAGGG	CCAGCCCAGA	ATGCCAGCCC	TCTTGTCCTT	1140
	CCGCCA						1146

- 20 (2) INFORMATION FOR SEQ ID NO: 20:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 951
 - (B) TYPE: Nucleic acid
 - (C) STRANDEDNESS: Double
- 25 (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: cDNA to mRNA
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- 30 (B) CELL KIND: Liver
 - (D) CLONE NAME: HP01299
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:
- 60 35 ATGTGGCTCT ACCTGGCGGC CTTCGTGGGC CTGTACTACC TTCTGCACTG GTACCGGGAG AGGCAGGTGG TGAGCCACCT CCAAGACAAG TATGTCTTTA TCACGGGCTG TGACTCGGGC 120 180 TTTGGGAACC TGCTGGCCAG ACAGCTGGAT GCACGAGGCT TGAGAGTGCT GGCTGCGTGT 240 CTGACGGAGA AGGGGGCCGA GCAGCTGAGG GGCCAGACGT CTGACAGGCT GGAGACGGTG

	ACCCTGGATG	TTACCAAGAT	GGAGAGCATC	GCTGCAGCTA	CTCAGTGGGT	GAAGGAGCAT	300
	GTGGGGGACA	GAGGACTCTG	GGGACTGGTG	AACAATGCAG	GCATTCTTAC	ACCAATTACC	360
	TTATGTGAGT	GGCTGAACAC	TGAGGACTCT	ATGAATATGC	TCAAAGTGAA	CCTCATTGGT	420
	GTGATCCAGG	TGACCTTGAG	CATGCTTCCT	TTGGTGAGGA	GAGCACGGGG	AAGAATTGTC	480
5	AATGTCTCCA	GCATTCTGGG	AAGAGTTGCT	TTCTTTGTAG	GAGGCTACTG	TGTCTCCAAG	540
	TATGGAGTGG	AAGCCTTTTC	AGATATTCTG	AGGCGTGAGA	TTCAACATTT	TGGGGTGAAA	600
	ATCAGCATAG	TTGAACCTGG	CTACTTCAGA	ACGGGAATGA	CAAACATGAC	ACAGTCCTTA	660
	GAGCGAATGA	AGCAAAGTTG	GAAAGAAGCC	CCCAAGCATA	TTAAGGAGAC	CTATGGACAG	720
	CAGTATTTTG	ATGCCCTTTA	CAATATCATG	AAGGAAGGC	TGTTGAATTG	TAGCACAAAC	780
.0	CTGAACCTGG	TCACTGACTG	CATGGAACAT	GCTCTGACAT	CGGTGCATCC	GCGAACTCGA	840
	TATTCAGCTG	GCTGGGATGC	TAAATTTTTC	TTCATCCCTC	TATCTTATTT	ACCTACATCA	900
	CTGGCAGACT	ACATTTTGAC	TAGATCTTGG	CCCAAACCAG	CCCAGGCAGT	С	951

15 (2) INFORMATION FOR SEQ ID NO: 21:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 888
 - (B) TYPE: Nucleic acid
 - (C) STRANDEDNESS: Double
- 20 (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: cDNA to mRNA

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- 25 (B) CELL KIND: Liver
 - (D) CLONE NAME: HP01347

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

30	ATGAGTGACT	CCAAGGAACC	AAGGGTGCAG	CAGCTGGGCC	TCCTGGGGTG	TCTTGGCCAT	60
	GGCGCCCTGG	TGCTGCAACT	CCTCTCCTTC	ATGCTCTTGG	CTGGGGTCCT	GGTGGCCATC	120
	CTTGTCCAAG	TGTCCAAGGT	CCCCAGCTCC	CTAAGTCAGG	AACAATCCGA	GCAAGACGCA	180
	ATCTACCAGA	ACCTGACCCA	GCTTAAAGCT	GCAGTGGGTG	AGCTCTCAGA	GAAATCCAAG	240
	CTGCAGGAGA	TCTACCAGGA	GCTGACCCAG	CTGAAGGCTG	CAGTGGGTGA	GTTGCCAGAG	300
35	AAATCCAAGC	TGCAGGAGAT	CTACCAGGAG	CTGACCCGGC	TGAAGGCTGC	AGTGGGTGAG	360
	TTGCCAGAGA	AATCCAAGCT	GCAGGAGATC	TACCAGGAGC	TGACCCGGCT	GAAGGCTGCA	420
	GTGGGTGAGT	TGCCAGAGAA	ATCCAAGCTG	CAGGAGATCT	ACCAGGAGCT	GACCCGGCTG	480
	AAGGCTGCAG	TGGGTGAGTT	GCCAGAGAAA	TCCAAGCTGC	AGGAGATCTA	CCAGGAGCTG	540

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	ACGGAGCTGA	AGGCTGCAGT	GGGTGAGTTG	CCAGAGAAAT	CCAAGCTGCA	GGAGATCTAC	600
	CAGGAGCTGA	CCCAGCTGAA	GGCTGCAGTG	GGTGAGTTGC	CAGACCAGTC	CAAGCAGCAG	660
	CAAATCTATC	AAGAACTGAC	CGATTTGAAG	ACTGCATTTG	AACGCCTGTG	CCGCCACTGT	720
	CCCAAGGACT	GGACATTCTT	CCAAGGAAAC	TGTTACTTCA	TGTCTAACTC	CCAGCGGAAC	780
5	TGGCACGACT	CCGTCACCGC	CTGCCAGGAA	GTGAGGGCCC	AGCTCGTCGT	AATCAAAACT	840
-			GGTACTGGAA				888

(2) INFORMATION FOR SEQ ID NO: 22:

10 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 591
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- 15 (ii) SEQUENCE KIND: cDNA to mRNA

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- 20 (D) CLONE NAME: HP01440

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

	ATGTGTACGG	GAAAATGTGC	CCGCTGTGTG	GGGCTCTCCC	TCATTACCCT	CTGCCTCGTC	60
25	TGCATTGTGG	CCAACGCCCT	CCTGCTGGTA	CCTAATGGGG	AGACCTCCTG	GACCAACACC	120
	AACCATCTCA	GCTTGCAAGT	CTGGCTCATG	GGCGGCTTCA	TTGGCGGGGG	CCTAATGGTA	180
	CTGTGTCCGG	GGATTGCAGC	CGTTCGGGCA	GGGGGCAAGG	GCTGCTGTGG	TGCTGGGTGC	240
	TGTGGAAACC	GCTGCAGGAT	GCTGCGCTCG	GTCTTCTCCT	CGGCGTTCGG	GGTGCTTGGT	300
	GCCATCTACT	GCCTCTCGGT	GTCTGGAGCT	GGGCTCCGAA	ATGGACCCAG	ATGCTTAATG	360
30	AACGGCGAGT	GGGGCTACCA	CTTCGAAGAC	ACCGCGGGAG	CTTACTTGCT	CAACCGCACT	420
	CTATGGGATC	GGTGCGAGGC	GCCCCTCGC	GTGGTCCCCT	GGAATGTGAC	GCTCTTCTCG	480
	CTGCTGGTGG	CCGCCTCCTG	CCTGGAGATA	GTACTGTGTG	GGATCCAGCT	GGTGAACGCG	540
			CGATTGCAGG				591

- (2) INFORMATION FOR SEQ ID NO: 23:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 663

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(B) TYPE: Nucleic acid(C) STRANDEDNESS: Double(D) TOPOLOGY: Linear

(ii) SEQUENCE KIND: cDNA to mRNA

5

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens(B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP01526

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

	ATGGAGGCGG	GCGGCTTTCT	GGACTCGCTC	ATTTACGGAG	CATGCGTGGT	CTTCACCCTT	60
	GGCATGTTCT	CCGCCGGCCT	CTCGGACCTC	AGGCACATGC	GAATGACCCG	GAGTGTGGAC	120
15	AACGTCCAGT	TCCTGCCCTT	TCTCACCACG	GAAGTCAACA	ACCTGGGCTG	GCTGAGTTAT	180
	GGGGCTTTGA	AGGGAGACGG	GATCCTCATC	GTCGTCAACA	CAGTGGGTGC	TGCGCTTCAG	240
	ACCCTGTATA	TCTTGGCATA	TCTGCATTAC	TGCCCTCGGA	AGCGTGTTGT	GCTCCTACAG	300
	ACTGCAACCC	TGCTAGGGGT	CCTTCTCCTG	GGTTATGGCT	ACTTTTGGCT	CCTGGTACCC	360
	AACCCTGAGG	CCCGGCTTCA	GCAGTTGGGC	CTCTTCTGCA	GTGTCTTCAC	CATCAGCATG	420
20	TACCTCTCAC	CACTGGCTGA	CTTGGCTAAG	GTGATTCAAA	CTAAATCAAC	CCAATGTCTC	480
	TCCTACCCAC	TCACCATTGC	TACCCTTCTC	ACCTCTGCCT	CCTGGTGCCT	CTATGGGTTT	540
	CGACTCAGAG	ATCCCTATAT	CATGGTGTCC	AACTTTCCAG	GAATCGTCAC	CAGCTTTATC	600
	CGCTTCTGGC	TTTTCTGGAA	GTACCCCCAG	GAGCAAGACA	GGAACTACTG	GCTCCTGCAA	660
	ACC .						663

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(2) INFORMATION FOR SEQ ID NO: 24:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 753
- 30 (B) TYPE: Nucleic acid
 - (C) STRANDEDNESS: Double
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: cDNA to mRNA

35 (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10230

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:

	ATGTCGGACA	TCGGAGACTG	GTTCAGGAGC	ATCCCGGCGA	TCACGCGCTA	TTGGTTCGCC	60
	GCCACCGTCG	CCGTGCCCTT	GGTCGGCAAA	CTCGGCCTCA	TCAGCCCGGC	CTACCTCTTC	120
5	CTCTGGCCCG	AAGCCTTCCT	TTATCGCTTT	CAGATTTGGA	GGCCAATCAC	TGCCACCTTT	180
	TATTTCCCTG	TGGGTCCAGG	AACTGGATTT	CTTTATTTGG	TCAATTTATA	TTTCTTATAT	240
	CAGTATTCTA	CGCGACTTGA	AACAGGAGCT	TTTGATGGGA	GGCCAGCAGA	CTATTTATTC	300
	ATGCTCCTCT	TTAACTGGAT	TTGCATCGTG	ATTACTGGCT	TAGCAATGGA	TATGCAGTTG	360
	CTGATGATTC	CTCTGATCAT	GTCAGTACTT	TATGTCTGGG	CCCAGCTGAA	CAGAGACATG	420
10	ATTGTATCAT	TTTGGTTTGG	AACACGATTT	AAGGCCTGCT	ATTTACCCTG	GGTTATCCTT	480
	GGATTCAACT	ATATCATCGG	AGGCTCGGTA	ATCAATGAGC	TTATTGGAAA	TCTGGTTGGA	540
	CATCTTTATT	TTTTCCTAAT	GTTCAGATAC	CCAATGGACT	TGGGAGGAAG	AAATTTTCTA	600
	TCCACACCTC	AGTTTTTGTA	CCGCTGGCTG	CCCAGTAGGA	GAGGAGGAGT	ATCAGGATTT	660
	GGTGTGCCCC	CTGCTAGCAT	GAGGCGAGCT	GCTGATCAGA	ATGGCGGAGG	CGGGAGACAC	720
15	AACTGGGGCC	AGGGCTTTCG	ACTTGGAGAC	CAG			753

(2) INFORMATION FOR SEQ ID NO: 25:

(i) SEQUENCE CHARACTERISTICS:

20 (A) LENGTH: 318

(B) TYPE: Nucleic acid

(C) STRANDEDNESS: Double

(D) TOPOLOGY: Linear

(ii) SEQUENCE KIND: cDNA to mRNA

25

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(B) CELL KIND: Epidermoid carcinoma

(C) CELL LINE: KB

30 (D) CLONE NAME: HP10389

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:

	ATGGCGACTC	CCGGCCCTGT	GATTCCGGAG	GTCCCCTTTG	AACCATCGAA	GCCTCCAGTC	60
35	ATTGAGGGGC	TGAGCCCCAC	TGTTTACAGG	AATCCAGAGA	GTTTCAAGGA	AAAGTTCGTT	120
	CGCAAGACCC	GCGAGAACCC	GGTGGTACCC	ATAGGTTGCC	TGGCCACGGC	GGCCGCCCTC	180
	ACCTACGGCC	TCTACTCCTT	CCACCGGGGC	AACAGCCAGC	GCTCTCAGCT	CATGATGCGC	240
	ACCCGGATCG	CCGCCCAGGG	TTTCACGGTC	GCAGCCATCT	TGCTGGGTCT	GGCTGTCACT	300

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GCTATGAAGT CTCGACCC

	(2) INFORMATION FOR SEQ ID NO: 26:	
5	(i) SEQUENCE CHARACTERISTICS:	
-	(A) LENGTH: 234	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
10	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Stomach cancer	
15	(D) CLONE NAME: HP10408	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:	
	ATGGGGTCTG GGCTGCCCCT TGTCCTCCTC TTGACCCTCC TTGGCAGCTC ACATGGAACA	60
20	GGGCCGGGTA TGACTTTGCA ACTGAAGCTG AAGGAGTCTT TTCTGACAAA TTCCTCCTAT	120
	GAGTCCAGCT TCCTGGAATT GCTTGAAAAG CTCTGCCTCC TCCTCCATCT CCCTTCAGGG	180
	ACCAGCGTCA CCCTCCACCA TGCAAGATCT CAACACCATG TTGTCTGCAA CACA	234
25	(2) INFORMATION FOR SEQ ID NO: 27:	
2.5	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 942	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
30	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
35	(B) CELL KIND: Stomach cancer	
	(D) CLONE NAME: HP10412	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

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	ATGGTGGCGC	CTGTGTGGTA	CTTGGTAGCG	GCGGCTCTGC	TAGTCGGCTT	TATCCTCTTC	60
	CTGACTCGCA	GCCGGGGCCG	GGCGGCATCA	GCCGGCCAAG	AGCCACTGCA	CAATGAGGAG	120
	CTGGCAGGAG	CAGGCCGGGT	GGCCCAGCCT	GGGCCCCTGG	AGCCTGAGGA	GCCGAGAGCT	180
	GGAGGCAGGC	CTCGGCGCCG	GAGGGACCTG	GGCAGCCGCC	TACAGGCCCA	GCGTCGAGCC	240
5	CAGCGGGTGG	CCTGGGCAGA	AGCAGATGAG	AACGAGGAGG	AAGCTGTCAT	CCTAGCCCAG	300
	GAGGAGGAAG	GTGTCGAGAA	GCCAGCGGAA	ACTCACCTGT	CGGGGAAAAT	TGGAGCTAAG	360
	AAACTGCGGA	AGCTGGAGGA	GAAACAAGCG	CGAAAGGCCC	AGCGTGAGGC	AGAGGAGGCT	420
	GAACGTGAGG	AGCGGAAACG	ACTCGAGTCC	CAGCGCGAAG	CTGAGTGGAA	GAAGGAGGAG	480
	GAGCGGCTTC	GCCTGGAGGA	GGAGCAGAAG	GAGGAGGAGG	AGAGGAAGGC	CCGCGAGGAG	540
10	CAGGCCCAGC	GGGAGCATGA	GGAGTACCTG	AAACTGAAGG	AGGCCTTTGT	GGTGGAGGAG	600
	GAAGGCGTAG	GAGAGACCAT	GACTGAGGAA	CAGTCCCAGA	GCTTCCTGAC	AGAGTTCATC	660
	AACTACATCA	AGCAGTCCAA	GGTTGTGCTC	TTGGAAGACC	TGGCTTCCCA	GGTGGGCCTA	720
	CGCACTCAGG	ACACCATAAA	TCGCATCCAG	GACCTGCTGG	CTGAGGGGAC	TATAACAGGT	780
	GTGATTGACG	ACCGGGGCAA	GTTCATCTAC	ATAACCCCAG	AGGAACTGGC	CGCCGTGGCC	840
15	AACTTCATCC	GACAGCGGGG	CCGGGTGTCC	ATCGCCGAGC	TTGCCCAAGC	CAGCAACTCC	900
	CTCATCGCCT	GGGGCCGGGA	GTCCCCTGCC	CAAGCCCCAG	CC		942

(2) INFORMATION FOR SEQ ID NO: 28:

20 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 585
- (B) TYPE: Nucleic acid
- (C) STRANDEDNESS: Double
- (D) TOPOLOGY: Linear
- 25 (ii) SEQUENCE KIND: cDNA to mRNA

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- 30 (D) CLONE NAME: HP10413

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

	ATGGCTGCCG	AGGATGTGGT	GGCGACTGGC	GCCGACCCAA	GCGATCTGGA	GAGCGGCGGG	60
35	CTGCTGCATG	AGATTTTCAC	GTCGCCGCTC	AACCTGCTGC	TGCTTGGCCT	CTGCATCTTC	120
	CTGCTCTACA	AGATCGTGCG	CGGGGACCAG	CCGGCGGCCA	GCGGCGACAG	CGACGACGAC	180
	GAGCCGCCCC	CTCTGCCCCG	CCTCAAGCGG	CGCGACTTCA	CCCCCGCCGA	GCTGCGGCGC	240
	TTCGACGGCG	TCCAGGACCC	GCGCATACTC	ATGGCCATCA	ACGGCAAGGT	GTTCGATGTG	300

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	ACCAAAGGCC	GCAAATTCTA	CGGGCCCGAG	GGGCCGTATG	GGGTCTTTGC	TGGAAGAGAT	360
	GCATCCAGGG	GCCTTGCCAC	ATTTTGCCTG	GATAAGGAAG	CACTGAAGGA	TGAGTACGAT	420
	GACCTTTCTG	ACCTCACTGC	TGCCCAGCAG	GAGACTCTGA	GTGACTGGGA	GTCTCAGTTC	480
	ACTTTCAAGT	ATCATCACGT	GGGCAAACTG	CTGAAGGAGG	GGGAGGAGCC	CACTGTGTAC	540
5	TCAGATGAGG	AAGAACCAAA	AGATGAGAGT	GCCCGGAAAA	ATGAT		585

(2) INFORMATION FOR SEQ ID NO: 29:

(i) SEQUENCE CHARACTERISTICS:

10 (A) LENGTH: 1386

(B) TYPE: Nucleic acid

(C) STRANDEDNESS: Double

(D) TOPOLOGY: Linear

(ii) SEQUENCE KIND: cDNA to mRNA

15

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(B) CELL KIND: Stomach cancer

(D) CLONE NAME: HP10415

20

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

	ATGTTGGACT	TCGCGATCTT	CGCCGTTACC	TTCTTGCTGG	CGTTGGTGGG	AGCCGTGCTC	60
	TACCTCTATC	CGGCTTCCAG	ACAAGCTGCA	GGAATTCCAG	GGATTACTCC	AACTGAAGAA	120
25	AAAGATGGTA	ATCTTCCAGA	TATTGTGAAT	AGTGGAAGTT	TGCATGAGTT	CCTGGTTAAT	180
	TTGCATGAGA	GATATGGGCC	TGTGGTCTCC	TTCTGGTTTG	GCAGGCGCCT	CGTGGTTAGT	240
	TTGGGCACTG	TTGATGTACT	GAAGCAGCAT	ATCAATCCCA	ATAAGACATT	GGACCCTTTT	300
	GAAACCATGC	TGAAGTCATT	ATTAAGGTAT	CAATCTGGTG	GTGGCAGTGT	GAGTGAAAAC	360
	CACATGAGGA	AAAATTGTA	TGAAAATGGT	GTGACTGATT	CTCTGAAGAG	TAACTTTGCC	420
30	CTCCTCCTAA	AGCTTTCAGA	AGAATTATTA	GATAAATGGC	TCTCCTACCC	AGAGACCCAG	480
	CACGTGCCCC	TCAGCCAGCA	TATGCTTGGT	TTTGCTATGA	AGTCTGTTAC	ACAGATGGTA	540
	ATGGGTAGTA	CATTTGAAGA	TGATCAGGAA	GTCATTCGCT	TCCAGAAGAA	TCATGGCACA	600
	GTTTGGTCTG	AGATTGGAAA	AGGCTTTCTA	GATGGGTCAC	TTGATAAAAA	CATGACTCGG	660
	AAAAAACAAT	ATGAAGATGC	CCTCATGCAA	CTGGAGTCTG	TTTTAAGGAA	CATCATAAAA	720
35	GAACGAAAAG	GAAGGAACTT	CAGTCAACAT	ATTTTCATTG	ACTCCTTAGT	ACAAGGGAAC	780
	CTTAATGACC	AACAGATCCT	AGAAGACAGT	ATGATATTT	CTCTGGCCAG	TTGCATAATA	840
	ACTGCAAAAT	TGTGTACCTG	GGCAATCTGT	TTTTTAACCA	CCTCTGAAGA	AGTTCAAAAA	900
	AAATTATATG	AAGAGATAAA	CCAAGTTTTT	GGAAATGGTC	CTGTTACTCC	AGAGAAAATT	960

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	GAGCAGCTCA	GATATTGTCA	GCATGTGCTT	TGTGAAACTG	TTCGAACTGC	CAAACTGACT	1020
	CCAGTTTCTG	CCCAGCTTCA	AGATATTGAA	GGAAAAATTG	ACCGATTTAT	TATTCCTAGA	1080
	GAGACCCTCG	TCCTTTATGC	CCTTGGTGTG	GTACTTCAGG	ATCCTAATAC	TTGGCCATCT	1140
	CCACACAAGT	TTGATCCAGA	TCGGTTTGAT	GATGAATTAG	TAATGAAAAC	TTTTTCCTCA	1200
5	CTTGGATTCT	CAGGCACACA	GGAGTGTCCA	GAGTTGAGGT	TTGCATATAT	GGTGACCACA	1260
	GTACTTCTTA	GTGTATTGGT	GAAGAGACTG	CACCTACTTT	CTGTGGAGGG	ACAGGTTATT	1320
	GAAACAAAGT	ATGAACTGGT	AACATCATCA	AGGGAAGAAG	CTTGGATCAC	TGTCTCAAAG	1380
	AGATAT						1386

10

(2) INFORMATION FOR SEQ ID NO: 30:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 741
 - (B) TYPE: Nucleic acid
- 15 (C) STRANDEDNESS: Double
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: cDNA to mRNA

(vi) ORIGINAL SOURCE:

20 (A) ORGANISM: Homo sapiens

(B) CELL KIND: Stomach cancer

(D) CLONE NAME: HP10419

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

25

30

ATGGGGGCTG	CGGTGTTTTT	CGGCTGCACT	TTCGTCGCGT	TCGGCCCGGC	CTTCGCGCTT	60
TTCTTGATCA	CTGTGGCTGG	GGACCCGCTT	CGCGTTATCA	TCCTGGTCGC	AGGGGCATTT	120
TTCTGGCTGG	TCTCCCTGCT	CCTGGCCTCT	GTGGTCTGGT	TCATCTTGGT	CCATGTGACC	180
GACCGGTCAG	ATGCCCGGCT	CCAGTACGGC	CTCCTGATTT	TTGGTGCTGC	TGTCTCTGTC	240
CTTCTACAGG	AGGTGTTCCG	CTTTGCCTAC	TACAAGCTGC	TTAAGAAGGC	AGATGAGGGG	300
TTAGCATCGC	TGAGTGAGGA	CGGAAGATCA	CCCATCTCCA	TCCGCCAGAT	GGCCTATGTT	360
TCTGGTCTCT	CCTTCGGTAT	CATCAGTGGT	GTCTTCTCTG	TTATCAATAT	TTTGGCTGAT	420
GCACTTGGGC	CAGGTGTGGT	TGGGATCCAT	GGAGACTCAC	CCTATTACTT	CCTGACTTCA	480
GCCTTTCTGA	CAGCAGCCAT	TATCCTGCTC	CATACCTTTT	GGGGAGTTGT	GTTCTTTGAT	540
GCCTGTGAGA	GGAGACGGTA	CTGGGCTTTG	GGCCTGGTGG	TTGGGAGTCA	CCTACTGACA	600
TCGGGACTGA	CATTCCTGAA	CCCCTGGTAT	GAGGCCAGCC	TGCTGCCCAT	CTATGCAGTC	660
ACTGTTTCCA	TGGGGCTCTG	GGCCTTCATC	ACAGCTGGAG	GGTCCCTCCG	AAGTATTCAG	720
CGCAGCCTCT	TGTGTAAGGA	С				741

PCT/JP98/02445 WO 98/55508

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	(2) INFORMATION FOR SEQ ID NO: 31:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 339	
5	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
10	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Stomach cancer	
	(D) CLONE NAME: HP10424	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:	
	ATGAACTTCT ATTTACTCCT AGCGAGCAGC ATTCTGTGTG CCTTGATTGT CTTCTGGAAA	60
	TATCGCCGCT TTCAGAGAAA CACTGGCGAA ATGTCATCAA ATTCAACTGC TCTTGCACTA	120
	GTGAGACCCT CTTCTTCTGG GTTAATTAAC AGCAATACAG ACAACAATCT TGCAGTCTAC	180
20	GACCTCTCTC GGGATATTTT AAATAATTTC CCACACTCAA TAGCCAGGCA GAAGCGAATA	240
	TTGGTAAACC TCAGTATGGT GGAAAACAAG CTGGTTGAAC TGGAACATAC TCTACTTAGC	300
	AAGGGTTTCA GAGGTGCATC ACCTCACCGG AAATCCACC	339
25	(2) INFORMATION FOR SEQ ID NO: 32:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1095	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
30	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	

(vi) ORIGINAL SOURCE:

35

(A) ORGANISM: Homo sapiens

(D) CLONE NAME: HP10428

(C) CELL LINE: KB

(B) CELL KIND: Epidermoid carcinoma

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:

	ATGGGGAGGT	${\tt GGGCCCTCGA}$	TGTGGCCTTT	TTGTGGAAGG	CGGTGTTGAC	CCTGGGGCTG	60
	GTGCTTCTCT	ACTACTGCTT	CTCCATCGGC	ATCACCTTCT	ACAACAAGTG	GCTGACAAAG	120
5	AGCTTCCATT	TCCCCCTCTT	CATGACGATG	CTGCACCTGG	CCGTGATCTT	CCTCTTCTCC	180
	GCCCTGTCCA	GGGCGCTGGT	TCAGTGCTCC	AGCCACAGGG	CCCGTGTGGT	GCTGAGCTGG	240
	GCCGACTACC	TCAGAAGAGT	GGCTCCCACA	GCTCTGGCGA	CGGCGCTTGA	CGTGGGCTTG	300
	TCCAACTGGA	GCTTCCTGTA	TGTCACCGTC	TCGCTGTACA	CAATGACCAA	ATCCTCAGCT	360
	GTCCTCTTCA	TCTTGATCTT	CTCTCTGATC	TTCAAGCTGG	AGGAGCTGCG	CGCGGCACTG	420
10	GTCCTGGTGG	TCCTCCTCAT	CGCCGGGGGT	CTCTTCATGT	TCACCTACAA	GTCCACACAG	480
	TTCAACGTGG	AGGGCTTCGC	CTTGGTGCTG	GGGGCCTCGT	TCATCGGTGG	CATTCGCTGG	540
	ACCCTCACCC	AGATGCTCCT	GCAGAAGGCT	GAACTCGGCC	TCCAGAATCC	CATCGACACC	600
	ATGTTCCACC	TGCAGCCACT	CATGTTCCTG	GGGCTCTTCC	CTCTCTTTGC	TGTATTTGAA	660
	GGTCTCCATT	TGTCCACATC	TGAGAAAATC	TTCCGTTTCC	AGGACACAGG	GCTGCTCCTG	720
15	CGGGTACTTG	GGAGCCTCTT	CCTTGGCGGG	ATTCTCGCCT	TTGGTTTGGG	CTTCTCTGAG	780
	TTCCTCCTGG	TCTCCAGAAC	CTCCAGCCTC	ACTCTCTCCA	TTGCCGGCAT	TTTTAAGGAA	840
	GTCTGCACTT	TGCTGTTGGC	AGCTCATCTG	CTGGGCGATC	AGATCAGCCT	CCTGAACTGG	900
	CTGGGCTTCG	CCCTCTGCCT	CTCGGGAATA	TCCCTCCACG	TTGCCCTCAA	AGCCCTGCAT	960
	TCCAGAGGTG	ATGGTGGCCC	CAAGGCCTTG	AAGGGGCTGG	GCTCCAGCCC	CGACCTGGAG	1020
20	CTGCTGCTCC	GGAGCAGCCA	GCGGGAGGAA	GGTGACAATG	AGGAGGAGGA	GTACTTTGTG	1080
	GCCCAGGGGC	AGCAG					1095

(2) INFORMATION FOR SEQ ID NO: 33:

- 25 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 678
 - (B) TYPE: Nucleic acid
 - (C) STRANDEDNESS: Double
 - (D) TOPOLOGY: Linear
- 30 (ii) SEQUENCE KIND: cDNA to mRNA

(vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- 35 (D) CLONE NAME: HP10429
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

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	11,	
	ATGCCTACCA CAAAGAAGAC ATTGATGTTC TTATCAAGCT TTTTCACCAG CCTTGGGTCC	60
	TTCATTGTAA TTTGCTCTAT TCTTGGGACA CAAGCATGGA TCACCAGTAC AATTGCTGTT	120
	AGAGACTCTG CTTCAAATGG GAGCATTTTC ATCACTTACG GACTTTTTCG TGGGGAGAGT	180
	AGTGAAGAAT TGAGTCACGG ACTTGCAGAA CCAAAGAAAA AGTTTGCAGT TTTAGAGATA	240
5	CTGAATAATT CTTCCCAAAA AACTCTGCAT TCGGTGACTA TCCTGTTCCT GGTCCTGAGT	300
	TTGATCACGT CGCTGCTGAG CTCTGGGTTT ACCTTCTACA ACAGCATCAG CAACCCTTAC	360
	CAGACATTCC TGGGGCCGAC GGGGGTGTAC ACCTGGAACG GGCTCGGTGC ATCCTTCGTT	420
	TTTGTGACCA TGATACTGTT TGTGGCGAAC ACGCAGTCCA ACCAACTCTC CGAAGAGTTG	480
	TTCCAAATGC TTTACCCGGC AACCACCAGT AAAGGAACGA CCCACAGTTA CGGATACTCG	540
10	TTCTGGCTCA TACTGCTCGT CATTCTTCTA AATATAGTCA CTGTAACCAT CATCATTTTC	600
	TACCAGAAGG CCAGATACCA GCGGAAGCAG GAGCAGAGAA AGCCAATGGA ATATGCTCCA	660
	AGGGACGGAA TTTTATTC	678
15	(2) INFORMATION FOR SEQ ID NO: 34:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 387	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
20	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
25	(B) CELL KIND: Liver	
	(D) CLONE NAME: HP10432	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:	
	•	
30		
	ATGGCTCGGG GCTCGCTGCG CCGGTTGCTG CGGCTCCTCG TGCTGGGGCT CTGGCTGG	60
	TTGCTGCGCT CCGTGGCCGG GGAGCAAGCG CCAGGCACCG CCCCTGCTC CCGCGGCAGC	120
	TCCTGGAGCG CGGACCTGGA CAAGTGCATG GACTGCGCGT CTTGCAGGGC GCGACCGCAC	180
	AGCGACTTCT GCCTGGGCTG CGCTGCAGCA CCTCCTGCCC CCTTCCGGCT GCTTTGGCCC	240
35	ATCCTTGGGG GCGCTCTGAG CCTGACCTTC GTGCTGGGGC TGCTTTCTGG CTTTTTGGTC	300

TGGAGACGAT GCCGCAGGAG AGAGAAGTTC ACCACCCCCA TAGAGGAGAC CGGCGGAGAG

GGCTGCCCAG CTGTGGCGCT GATCCAG

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	(2) INFORMATION FOR SEQ ID NO: 35:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 489	
	(B) TYPE: Nucleic acid	
5	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
LO	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Liver	
	(D) CLONE NAME: HP10433	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:	
L5		
	ATGCGACGGC TGCTGATCCC TCTGGCCCTG TGGCTGGGCG CGGTGGGCGT GGGCGTCGCC	60
	GAGCTCACGG AAGCCCAGCG CCGGGGCCTG CAGGTGGCCC TGGAGGAATT TCACAAGCAC	120
	CCGCCCGTGC AGTGGGCCTT CCAGGAGACC AGTGTGGAGA GCGCCGTGGA CACGCCCTTC	180
	CCAGCTGGAA TATTTGTGAG GCTGGAATTT AAGCTGCAGC AGACAAGCTG CCGGAAGAGG	240
20	GACTGGAAGA AACCCGAGTG CAAAGTCAGG CCCAATGGGA GGAAACGGAA ATGCCTGGCC	300
	TGCATCAAAC TGGGCTCTGA GGACAAAGTT CTGGGCCGGT TGGTCCACTG CCCCATAGAG	360
	ACCCAAGTTC TGCGGGAGGC TGAGGAGCAC CAGGAGACCC AGTGCCTCAG GGTGCAGCGG	420
	GCTGGTGAGG ACCCCCACAG CTTCTACTTC CCTGGACAGT TCGCCTTCTC CAAGGCCCTG	480
	CCCCGCAGC	489
25		
	(2) INFORMATION FOR SEQ ID NO: 36:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 579	
30	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(22) 02(02)02 2200 00 1000	
35	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	

(B) CELL KIND: Stomach cancer

(D) CLONE NAME: HP10480

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

	ATGATCCGCT GCGGCCTGGC CTGCGAGCGC TGCCGCTGGA TCCTGCCCCT GCTCCTACTC	60
	AGCGCCATCG CCTTCGACAT CATCGCGCTG GCCGGCCGCG GCTGGTTGCA GTCTAGCGAC	120
5	CACGGCCAGA CGTCCTCGCT GTGGTGGAAA TGCTCCCAAG AGGGCGGCGG CAGCGGGTCC	180
	TACGAGGAGG GCTGTCAGAG CCTCATGGAG TACGCGTGGG GTAGAGCAGC GGCTGCCATG	240
	CTCTTCTGTG GCTTCATCAT CCTGGTGATC TGTTTCATCC TCTCCTTCTT CGCCCTCTGT	300
	GGACCCCAGA TGCTTGTCTT CCTGAGAGTG ATTGGAGGTC TCCTTGCCTT GGCTGCTGTG	360
	TTCCAGATCA TCTCCCTGGT AATTTACCCC GTGAAGTACA CCCAGACCTT CACCCTTCAT	420
10	GCCAACCGTG CTGTCACTTA CATCTATAAC TGGGCCTACG GCTTTGGGTG GGCAGCCACG	480
	ATTATCCTGA TCGGCTGTGC CTTCTTCTTC TGCTGCCTCC CCAACTACGA AGATGACCTT	540
	CTGGGCAATG CCAAGCCCAG GTACTTCTAC ACATCTGCC	579
15	(2) INFORMATION FOR SEQ ID NO: 37:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1502	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
20	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
25	(B) CELL KIND: Liver	
	(D) CLONE NAME: HP01263	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
30	(B) EXISTENCE POSITION: 37 1185	
30	(C) CHARACTERIZATION METHOD: E	
	(b) Characterization introd. 1	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:	
	(XI) SEQUENCE DESCRIPTION: SEQ ID NO. 37.	
35	ACAAACTGAC CCATCCTGGG CCTTGTTCTC CACAGA ATG GGT CTG CTC CTT CCC	54
55	Met Gly Leu Leu Pro	
	1 5	
	CTG GCA CTC TGC ATC CTA GTC CTG TGC TGC GGA GCA ATG TCT CCA CCC	102
	010 000 010 100 010 010 010 100 000 000	

	Leu	Ala	Leu	Cys	Ile	Leu	Val	Leu	Cys	Cys	Gly	Ala	Met	Ser	Pro	Pro	
				10					15					20			
	CAG	CTG	GCC	CTC	AAC	CCC	TCG	GCT	CTG	CTC	TCC	ÇGG	GGC	TGC	AAT	GAC	150
	Gln	Leu	Ala	Leu	Asn	Pro	Ser	Ala	Leu	Leu	Ser	Arg	Gly	Cys	Asn	Asp	
5			25					30					35				
	TCC	GAT	GTG	CTG	GCA	GTT	GCA	GGC	TTT	GCC	CTG	CGG	GAT	ATT	AAC	AAA	198
	Ser	Asp	Val	Leu	Ala	Val	Ala	Gly	Phe	Ala	Leu	Arg	Asp	Ile	Asn	Lys	
		40					45					50					
	GAC	AGA	AAG	GAT	GGC	TAT	GTG	CTG	AGA	CTC	AAC	CGA	GTG	AAC	GAC	GCC	246
LO	Asp	Arg	Lys	Asp	Gly	Tyr	Val	Leu	Arg	Leu	Asn	Arg	Va1	Asn	Asp	Ala	
	55					60					65					70	
	CAG	GAA	TAC	AGA	CGG	GGT	GGC	CTG	GGA	TCT	CTG	TTC	TAT	CTT	ACA	CTG	294
	Gln	Glu	Tyr	Arg	Arg	Gly	Gly	Leu	Gly	Ser	Leu	Phe	Tyr	Leu	Thr	Leu	
					75					80					85		
15							TGC										342
	Asp	Val	Leu	Glu	Thr	Asp	Cys	His	Va1	Leu	Arg	Lys	Lys	Ala	Trp	Gln	
				90					95					100			
							TTT										390
	Asp	Cys	Gly	Met	Arg	Ile	Phe	Phe	Glu	Ser	Val	Tyr	Gly	Gln	Cys	Lys	
20			105					110					115				
							AAC										438
	Ala	Ile	Phe	Tyr	Met	Asn	Asn	Pro	Ser	Arg	Val		Tyr	Leu	Ala	Ala	
		120					125					130				400	400
							CCA										486
25		Asn	Cys	Thr	Leu		Pro	Val	Ser	Lys		Lys	TIE	Tyr	met		
	135					140					145				010	150	E 2 /
							TCC										534
	Cys	Pro	Asp	Cys		Ser	Ser	Ile	Pro		Asp	Ser	Ser	Asn		GIN	
					155					160					165	A A C	507
30							GAG										582
	Val	Leu	Glu		Ala	Thr	Glu	Ser		Ala	Lys	Tyr	Asn		GIU	Asn	
				170					175				000	180	400	CAC	620
							CTC										630
	Thr	Ser			Tyr	Ser	Leu		Lys	Val	Thr	Arg		Ser	ser	GIN	
35			185					190					195		244	mc A	676
																TCA	678
	Trp			Gly	Pro	Ser	Tyr		Val	Glu	Tyr		Ile	Lys	Glu	ser	
		200					205					210					

CCA TGT ACT AAA TCC CAG GCC AGC AGC TGT TCA CTT CAG TCC TCC GAC Pro Cys Thr Lys Ser Gln Ala Ser Ser Cys Ser Leu Gln Ser Ser Asp TCT GTG CCT GTT GGT CTT TGC AAA GGT TCT CTG ACT CGA ACA CAC TGG Ser Val Pro Val Gly Leu Cys Lys Gly Ser Leu Thr Arg Thr His Trp GAA AAG TTT GTC TCT GTG ACT TGT GAC TTC TTT GAA TCA CAG GCT CCA Glu Lys Phe Val Ser Val Thr Cys Asp Phe Phe Glu Ser Gln Ala Pro 10 GCC ACT GGA AGT GAA AAC TCT GCT GTT AAC CAG AAA CCT ACA AAC CTT Ala Thr Gly Ser Glu Asn Ser Ala Val Asn Gln Lys Pro Thr Asn Leu CCC AAG GTG GAA GAA TCC CAG CAG AAA AAC ACC CCC CCA ACA GAC TCC Pro Lys Val Glu Glu Ser Gln Gln Lys Asn Thr Pro Pro Thr Asp Ser CCC TCC AAA GCT GGG CCA AGA GGA TCT GTC CAA TAT CTT CCT GAC TTG Pro Ser Lys Ala Gly Pro Arg Gly Ser Val Gln Tyr Leu Pro Asp Leu GAT GAT AAA AAT TCC CAG GAA AAG GGC CCT CAG GAG GCC TTT CCT GTG 20 Asp Asp Lys Asn Ser Gln Glu Lys Gly Pro Gln Glu Ala Phe Pro Val CAT CTG GAC CTA ACC ACG AAT CCC CAG GGA GAA ACC CTG GAT ATT TCC His Leu Asp Leu Thr Thr Asn Pro Gln Gly Glu Thr Leu Asp Ile Ser 25 TTC CTC TTC CTG GAG CCT ATG GAG GAG AAG CTG GTT GTC CTG CCT TTC Phe Leu Phe Leu Glu Pro Met Glu Glu Lys Leu Val Val Leu Pro Phe CCC AAA GAA AAA GCA CGC ACT GCT GAG TGC CCA GGG CCA GCC CAG AAT Pro Lys Glu Lys Ala Arg Thr Ala Glu Cys Pro Gly Pro Ala Gln Asn GCC AGC CCT CTT GTC CTT CCG CCA TGAGAATCAC ACAGAGTCTT CTGTAGGG Ala Ser Pro Leu Val Leu Pro Pro GTATGGTGCG CCGCATGACA TGGGAGGCGA TGGGGACGAT GGACAGAGAC AGAGCGTGCA 35 CACGTAGAGT GGCTAGTGAA GGACGCCTTT TTGACTCTTC TTGGTCTCAG CATGTTGACT GGGATTGGAA ATAATGAGAC TGAGCCCTCG GCTTGGGCTG CACTCTACCC TGTACACTGC CTTGTACCCT GAGCTGCATC ACCTCCTAAA CTGAGCAGTC TCATACCATG GAGAGATGCC TCTCTTATGT CTTCAGCCAC TCACTTATAA AGATACTTAT CTTTTCAGCA GT

	(2)	INFO)KMA:	LION	FOR	SEQ	ו ענו	NO: .	38:								
		(i	i) SI	EQUE	NCE C	CHARA	ACTE	RIST	ics:								
				(A)	LENG	STH:	1349	•									
5				(B)	TYPE	E: Nu	ıclei	ic a	cid								
				(C)	STRA	ANDEI	ONESS	5: Do	ouble	2							
				(D)	TOPO	DLOGY	7: Li	inear	r								
		(5	ii) :	SEQUI	ENCE	KINI	o: cI	ONA 1	to mI	ANS							
LO		(1	/i) (ORIG	INAL	SOUT	RCE:										
				(A)	ORGA	ANISM	4: H	ото .	sapi	ens							
				(B)	CELI	L KI	VD: I	Live	r								
				(D)	CLO	NE NA	AME:	нро	1299								
15																	
		(:	ix) :	SEQUI	ENCE	CHAI	RACTI	ERIS	rics:	:							
				(A)	CHAI	RACTI	ERIZA	ATIO	N COI	DE: 0	CDS						
				(B)	EXIS	STEN	CE PO	OSIT:	ION:	111	10	064					
				(C)	CHAI	RACTI	ERIZA	OITA	N ME	HOD:	: E						
20																	
		(:	xi)	SEQUI	ENCE	DESC	CRIP:	rion	: SEC) ID	NO:	38:					
	AGCA	AGTTO	GGG (GCAG	GAGG	AA GO	CCGA	CTGC:	r gco	CTGG	CTG	CAAA	AGAA(GTC (CTTT	CAAGTC	60
									C GAO								116
25															Met !		
															1		
	CTC	TAC	CTG	GCG	GCC	TTC	GTG	GGC	CTG	TAC	TAC	CTT	CTG	CAC	TGG	TAC	
	164																
	Leu	Tyr	Leu	Ala	Ala	Phe	Val	Gly	Leu	Tyr	Tyr	Leu	Leu	His	Trp	Tyr	
30			5					10					15				
	CGG	GAG	AGG	CAG	GTG	GTG	AGC	CAC	CTC	CAA	GAC	AAG	TAT	GTC	TTT	ATC	212
	Arg	Glu	Arg	Gln	Val	Val	Ser	His	Leu	Gln	Asp	Lys	Tyr	Val	Phe	lle	
		20					25					30					
	ACG	GGC	TGT	GAC	TCG	GGC	TTT	GGG	AAC	CTG	CTG	GCC	AGA	CAG	CTG	GAT	260
35	Thr	Gly	Cys	Asp	Ser	Gly	Phe	Gly	Asn	Leu	Leu	Ala	Arg	Gln	Leu	Asp	
	35					40					45					50	
	GCA	CGA	GGC	TTG	AGA	GTG	CTG	GCT	GCG	TGT	CTG	ACG	GAG	AAG	GGG	GCC	308
	41.	4=~	C1	1 011	۸	Vo 1	1 011	۸1.	A 1 a	Cve	1 on	Th r	Glu	1.00	Glv	Ala	

					55					60					65		
	GAG	CAG	CTG	AGG	GGC	CAG	ACG	TCT	GAC	AGG	CTG	GAG	ACG	GTG	ACC	CTG	356
	Glu	Gln	Leu	Arg	Gly	Gln	Thr	Ser	Asp	Arg	Leu	Glu	Thr	Val	Thr	Leu	
				70					75					80			
5	GAT	GTT	ACC	AAG	ATG	GAG	AGC	ATC	GCT	GCA	GCT	ACT	CAG	TGG	GTG	AAG	404
	Asp	Val	Thr	Lys	Met	Glu	Ser	Ile	Ala	Ala	Ala	Thr	Gln	Trp	Val	Lys	
			85					90					95				
	GAG	CAT	GTG	GGG	GAC	AGA	GGA	CTC	TGG	GGA	CTG	GTG	AAC	AAT	GCA	GGC	452
	Glu	His	Val	Gly	Asp	Arg	Gly	Leu	Trp	Gly	Leu	Val	Asn	Asn	Ala	Gly	
10		100					105					110					
	ATT	CTT	ACA	CCA	ATT	ACC	TTA	TGT	GAG	TGG	CTG	AAC	ACT	GAG	GAC	TCT	500
	Ile	Leu	Thr	Pro	Ile	Thr	Leu	Cys	Glu	Trp	Leu	Asn	Thr	Glu	Asp	Ser	
	115					120					125					130	
	ATG	AAT	ATG	CTC	AAA	GTG	AAC	CTC	ATT	GGT	GTG	ATC	CAG	GTG	ACC	TTG	548
15	Met	Asn	Met	Leu	Lys	Val	Asn	Leu	Ile	Gly	Val	Ile	Gln	Val	Thr	Leu	
					135					140					145		
	AGC	ATG	CTT	CCT	TTG	GTG	AGG	AGA	GCA	CGG	GGA	AGA	ATT	GTC	AAT	GTC	596
	Ser	Met	Leu	Pro	Leu	Val	Arg	Arg	Ala	Arg	Gly	Arg	Ile	Val	Asn	Val	
				150					155					160			
20	TCC	AGC	ATT	CTG	GGA	AGA	GTT	GCT	TTC	TTT	GTA	GGA	GGC	TAC	TGT	GTC	644
	Ser	Ser	Ile	Leu	Gly	Arg	Val	Ala	Phe	Phe	Va1	Gly	Gly	Tyr	Cys	Val	
			165					170					175				
	TCC	AAG	TAT	GGA	GTG	GAA	GCC	TTT	TCA	GAT	ATT	CTG	AGG	CGT	GAG	ATT	692
	Ser	Lys	Tyr	Gly	Val	Glu	Ala	Phe	Ser	Asp	Ile	Leu	Arg	Arg	Glu	Ile	
25		180					185					190					
	CAA	CAT	TTT	GGG	ĢTG	AAA	ATC	AGC	ATA	GTT	GAA	CCT	GGC	TAC	TTC	AGA	740
	Gln	His	Phe	Gly	Val	Lys	Ile	Ser	Ile	Val	Glu	Pro	Gly	Tyr	Phe	Arg	
	195					200					205					210	
	ACG	GGA	ATG	ACA	AAC	ATG	ACA	CAG	TCC	TTA	GAG	CGA	ATG	AAG	CAA	AGT	788
30	Thr	Gly	Met	Thr	Asn	Met	Thr	Gln	Ser	Leu	Glu	Arg	Met	Lys	Gln	Ser	
					215					220					225		
	TGG	AAA	GAA	GCC	CCC	AAG	CAT	ATT	AAG	GAG	ACC	TAT	GGA	CAG	CAG	TAT	836
	Trp	Lys	Glu	Ala	Pro	Lys	His	Ile	Lys	Glu	Thr	Tyr	Gly	Gln	Gln	Tyr	
				230					235					240			
35	TTT	GAT	GCC	CTT	TAC	AAT	ATC	ATG	AAG	GAA	GGG	CTG	TTG	TAA	TGT	AGC	884
	Phe	Asp	Ala	Leu	Tyr	Asn	Ile	Met	Lys	Glu	Gly	Leu	Leu	Asn	Cys	Ser	
			245					250					255				
	ACA	AAC	CTG	AAC	CTG	GTC	ACT	GAC	TGC	ATG	GAA	CAT	GCT	CTG	ACA	TCG	932

124

	Thr A	sn Leu	Asn	Leu	Val	Thr	Asp	Cys	Met	Glu	His	Ala	Leu	Thr	Ser	
	2	60				265					270					
	GTG C	AT CCG	CGA	ACT	CGA	TAT	TCA	GCT	GGC	TGG	GAT	GCT	AAA	TTT	TTC	980
	Val H	is Pro	Arg	Thr	Arg	Tyr	Ser	Ala	Gly	Trp	Asp	Ala	Lys	Phe	Phe	
5	275				280					285					290	
	TTC A	TC CCT	CTA	TCT	TAT	TTA	CCT	ACA	TCA	CTG	GCA	GAC	TAC	ATT	TTG	1028
	Phe I	le Pro	Leu	Ser	Tyr	Leu	Pro	Thr	Ser	Leu	Ala	Asp	Tyr	Ile	Leu	
				295					300					305		
	ACT A	GA TCT	TGG	CCC	AAA	CCA	GCC	CAG	GCA	GTC	TAA	AGAA	AAC	TGGG	TTGGT	1080
10	Thr A	rg Ser	Trp	Pro	Lys	Pro	Ala	Gln	Ala	Val						
•			310					315								
	GCTTC	TTGGA .	ATGA	AGGC	AA AA	AATC'	TGAA	A TTO	GTTA(GTGT	CTC	AGTA	ATC	CTGA'	TTTAGA	1140
	ACCCA	GGCTT	TTTG'	raac.	AA T	GTGT:	TTTC	r TG	CCTA	TTAA	CAT	TAT	CTG	GCAT	CATCAG	1200
	AGTAC	TAACA	TGTT'	'ATAI	rr ro	CAGA'	TATC	C AA	AGCT'	TACC	ACT	TTAG	GTG .	ATGA	ATCTTT	1260
15	ACTAT	TTTAG	CCCT	rttt:	rg A	TGAG.	ACTA	r TT	GTCTA	AAAG	TGA	ATCA	TTT	GTTC	TTGCCT	1320
	TATTA	AACAG .	AGTA	GATG	GA A	AACA	ATTT									1349
	(2) I	NFORMA	TION	FOR	SEQ	ID 1	NO: :	39:								
20		(i) S	EQUE	NCE (CHAR	ACTE	RIST:	ics:								
			(A)	LEN	GTH:	164	3									
			(B)	TYP	E: N	ucle	ic a	cid								
			• •				S: Do		е							
			• •				inear									
25		(ii)	SEQU	ENCE	KIN	D: cl	DNA 1	to mi	RNA							
		(vi)														
							omo .	-	ens							
							Live									
30			(D)	CLO	NE NA	AME:	HPO:	1347								
		43.3	0.0011	-NO-	CILA			7700								
		(ix)	-							SDC.						
							ATIO				-					
			(B)	EXI	STEN(CE P	OSIT:	TON:	25.	. 91:)					

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

(C) CHARACTERIZATION METHOD: E

	AACA	TCT	GG (GACAG	CGGG	SA AA											51
							ì		Ser .	Asp S	Ser :	Lys C	lu I	Pro A	Arg \	/aı	
	CAC	CVC	CTC	GGC	ር ፓር	ርሞር	GGG	1 TGT	ር ጥጥ	GGC	ር ል T	5 eec	GCC	CTG	GTG	CTG	99
E				Gly													
5		GIN	Leu	GLY	Leu	15	GIY	Cys	ren	GIY	20	Gry	VTG	шси	***	25	
	10	CTC.	CTC	TCC	ጥጥር		ርጥሮ	ጥጥ ር	CCT	GGG		ርሞር	GTG.	GCC	ATC		147
				Ser													
	GIN	Leu	Leu	ser	30	Mec	Leu	Ten	nia	.35	Val	Deu	V 11.	7114	40	200	
. ^	O TO C	C A A	CTC	TCC		ርሞር	ccc	۸۵۲	ሞርር		ልር ጥ	CAG	CAA	CAA		GAG	195
10				Ser													
	Vai	GIII	val	45	Lys	Vai	FIU	361	50	Deu	JCL	0111	0.44	55	001		
	CAA	GAC	GCA	ATC	TAC	CAG	AAC	CTG		CAG	стт	AAA	GCT		GTG	GGT	243
				Ile													
15	GIII	nap	60	110	1,1	0111	11511	65		01	200	2,0	70			,	
1.5	GAG	СТС		GAG	AAA	TCC	AAG		CAG	GAG	ATC	TAC		GAG	CTG	ACC	291
				Glu													
	014	75			_,,		80					85					
	CAG		AAG	GCT	GCA	GTG		GAG	TTG	CCA	GAG	AAA	TCC	AAG	CTG	CAG	339
20				Ala													
	90		,			95	•				100					105	
	GAG	ATC	TAC	CAG	GAG	CTG	ACC	CGG	CTG	AAG	GCT	GCA	GTG	GGT	GAG	TTG	387
	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	Lys	Ala	-Ala	Val	Gly	Glu	Leu	
					110					115					120		
25	CCA	GAG	AAA	TCC	AAG	CTG	CAG	GAG	ATC	TAC	CAG	GAG	CTG	ACC	CGG	CTG	435
	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	
				125					130					135			
	AAG	GCT	GCA	GTG	GGT	GAG	TTG	CCA	GAG	AAA	TCC	AAG	CTG	CAG	GAG	ATC	483
	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	
30			140					145					150				
	TAC	CAG	GAG	CTG	ACC	CGG	CTG	AAG	GCT	GCA	GTG	GGT	GAG	TTG	CCA	GAG	531
	Tyr	Gln	Glu	Leu	Thr	Arg	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Glu	
		155					160					165					
	AAA	TCC	AAG	CTG	CAG	GAG	ATC	TAC	CAG	GAG	CTG	ACG	GAG	CTG	AAG	GCT	579
35	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	Glu	Leu	Thr	Glu	Leu	Lys	Ala	
	170					175					180)				185	
	GCA	GTG	GGT	GAG	TTG	CCA	GAG	AAA	TCC	AAG	CTG	CAG	GAG	ATC	TAC	CAG	627
	Ala	Val	Gly	Glu	Leu	Pro	Glu	Lys	Ser	Lys	Leu	Gln	Glu	Ile	Tyr	Gln	

					190					195					200		
	GAG	CTG	ACC	CAG	CTG	AAG	GCT	GCA	GTG	GGT	GAG	TTG	CCA	GAC	CAG	TCC	675
	Glu	Leu	Thr	Gln	Leu	Lys	Ala	Ala	Val	Gly	Glu	Leu	Pro	Asp	Gln	Ser	
				205					210					215			
5	AAG	CAG	CAG	CAA	ATC	TAT	CAA	GAA	CTG	ACC	GAT	TTG	AAG	ACT	GCA	TTT	723
	Lys	Gln	Gln	Gln	Ile	Tyr	Gln	Glu	Leu	Thr	Asp	Leu	Lys	Thr	Ala	Phe	
			220					225					230				
	GAA	CGC	CTG	TGC	CGC	CAC	TGT	ccc	AAG	GAC	TGG	ACA	TTC	TTC	CAA	GGA	771
	Glu	Arg	Leu	Cys	Arg	His	Cys	Pro	Lys	Asp	Trp	Thr	Phe	Phe	Gln	Gly	
10		235				•	240					245					
	AAC	TGT	TAC	TTC	ATG	TCT	AAC	TCC	CAG	CGG	AAC	TGG	CAC	GAC	TCC	GTC	819
	Asn	Cys	Tyr	Phe	Met	Ser	Asn	Ser	Gln	Arg	Asn	Trp	His	Asp	Ser	Val	
	250					255					260					265	
	ACC	GCC	TGC	CAG	GAA	GTG	AGG	GCC	CAG	CTC	GTC	GTA	ATC	AAA	ACT	GCT	867
15	Thr	Ala	Cys	Gln	Glu	Val	Arg	Ala	Gln	Leu	Val	Val	Ile	Lys	Thr	Ala	
					270					275					280		
	GAG	GAG	CAG	CTT	CCA	GCG	GTA	CTG	GAA	CAG	TGG	AGA	ACC	CAA	CAA		912
	Glu	Glu	Gln	Leu	Pro	Ala	Val	Leu	Glu	Gln	Trp	Arg	Thr	Gln	Gln		
				285					290					295			
20															ATCG		970
																TAGTTG	
																AAGCCA	
																TTCGAT	
																TCCCTG	
25																GGATGC	
																GATTAG	
				-												GCAAAC	
																GCTGGT	
															4	ACCAAT	
30																TCTTTG	
					TCCA	тт т	GGCT	GTTT	C TG.	AGTT(GTAG	CCT	TTAT	AAI	MAAG	TGGTAA	
	ATG'	TTGT.	AAC	TGC													1643

35 (2) INFORMATION FOR SEQ ID NO: 40:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 729

(B) TYPE: Nucleic acid

				(C)	STRA	NDED	NESS	: Do	uble	!							
				(D)	TOPO	LOGY	: Li	near	•								
		(i	.i) S	EQUE	NCE	KIND	: cD	NA t	o mR	NA							
5		(v	i) C	RIGI	NAL	SOUR	.CE:										
				(A)	ORGA	NISM	1: Hc	mo s	sapie	ens							
				(B)	CELL	. KIN	D: S	toma	ch c	ance	r						
				(D)	CLON	IE NA	ME:	HP01	440								
10		i)	(x) S	EQUE													
				(A)	CHAR	RACTE	RIZA	MOIT	1 COI	E: C	DS						
				(B)	EXIS	STENC	E PC	SITI	ON:	38	631	L					
				(C)	CHAF	RACTE	ERIZA	MOIT	ME1	HOD:	E						
15		()	(i) S	SEQUE	ENCE	DESC	RIPT	: 1ON:	: SEC	5 TD	NO:	40:					
	4 C TO	3mo 4 6	.mc /		~~~~	PC C9	יייירייי	7C A C /		PC & C (• ለጥ <i>ር</i>	י יייביי	ኮ ልሮር	2 66/	A A A A	A TGT	55
	ACT	TUAC	JIC F	ACCGC	.6161		1166	GAC	1 001	CACC						s Cys	
											1100	•			•	5	
20	GCC	CGC	TGT	GTG	GGG	CTC	TCC	CTC	ATT	ACC	_	_	CTC	GTC	TGC	ATT	103
20				Val													
		6		10	,				15			•		20			
	GTG	GCC		GCC	CTC	CTG	CTG	GTA	CCT	AAT	GGG	GAG	ACC	TCC	TGG	ACC	151
				Ala													
25			25					30					35				
	AAC	ACC	AAC	CAT	CTC	AGC	TTG	CAA	GTC	TGG	CTC	ATG	GGC	GGC	TTC	ATT	199
	Asn	Thr	Asn	His	Leu	Ser	Leu	Gln	Val	Trp	Leu	Met	Gly	Gly	Phe	Ile	
		40					45					50					
	GGC	GGG	GGC	CTA	ATG	GTA	CTG	TGT	CCG	GGG	ATT	GCA	GCC	GTT	CGG	GCA	247
30	Gly	Gly	Gly	Leu	Met	Val	Leu	Cys	Pro	Gly	Ile	Ala	Ala	Val	Arg	Ala	
	55					60					65					70	
				GGC													295
	Gly	Gly	Lys	Gly	Cys	Cys	Gly	Ala	Gly	Cys	Cys	Gly	Asn	Arg	Cys	Arg	
					75					80					85		
35				TCG													343
	Met	Leu	Arg	Ser	Val	Phe	Ser	Ser	Ala	Phe	Gly	Val	Leu		Ala	Ile	
				90					95					100			
	ጥለብ	TCC	CTC	TCC	CTC	ጥርጥ	CCA	COT	CCC	CTC	CGA	ΔΔΤ	CCA	CCC	AGA	TGC	391

	Tyr	Cys	Leu	Ser	Val	Ser	Gly	Ala	G1y	Leu	Arg	Asn	Gly	Pro	Arg	Cys	
			105					110					115				
	ATT	ATG	AAC	GGC	GAG	TGG	GGC	TAC	CAC	TTC	GAA	GAC	ACC	GCG	GGA	GCT	439
	Leu	Met	Asn	Gly	Glu	Trp	Gly	Tyr	His	Phe	Glu	Asp	Thr	Ala	Gly	Ala	
5		120	•				125					130					
	TAC	TTG	CTC	AAC	CGC	ACT	CTA	TGG	GAT	CGG	TGC	GAG	GCG	CCC	CCT	CGC	487
	Tyr	Leu	Leu	Asn	Arg	Thr	Leu	Trp	Asp	Arg	Cys	Glu	Ala	Pro	Pro	Arg	
	135					140					145					150	
	GTG	GTC	ccc	TGG	AAT	GTG	ACG	CTC	TTC	TCG	CTG	CTG	GTG	GCC	GCC	TCC	535
10	Val	Val	Pro	Trp	Asn	Val	Thr	Leu	Phe	Ser	Leu	Leu	Val	Ala	Ala	Ser	
					155					160					165		
	TGC	CTG	GAG	ATA	GTA	CTG	TGT	GGG	ATC	CAG	CTG	GTG	AAC	GCG	ACC	ATT	583
	Cys	Leu	Glu	Ile	Va1	Leu	Cys	Gly	Ile	Gln	Leu	Val	Asn	Ala	Thr	Ile	
			_	170					175					180			
15	GGT	GTC	TTC	TGC	GGC	GAT	TGC	AGG	AAA	AAA	CAG	GAC	ACC	CCT	CAC	TG	630
	Gly	Val	Phe	Cys	Gly	Asp	Cys	Arg	Lys	Lys	Gln	Asp	Thr	Pro	His		
			185					190					195				
	AGG	CTCC	ACT (GACC	GCCG	GG T	TACA	CTG	C TC	CTTC	CTGG	ACG	CCTA	CCT	GCT	CGCTCA	690
	CTC	CTT	GCT (CGCT	AGAA'	TA A	ACTG	CTTT	G CG	CTCT	CTT						729
20																	
	(2)						ID I										
		(:	i) S	•			ACTE		ICS:								
							132										
25							ucle.										
							DNES			е							
			• • • •				Y: L			D 17 A							
		(:	11) :	SEQU.	ENCE	KIN.	D: c	DNA '	to m	RNA							
٦.				0 D T O	T	0075	DOT .										
30		C	V1) (INAL												
							M: <i>H</i>		-								
							ND:			canc	er						
				(D)	CLO.	NE N.	AME:	HPU.	1326								
35		(ix)	SEQU	ENCE	CHA	RACT	ERIS	TICS	:							
		·		-			ERIZ.				CDS						
				•													
				(B)	EXI	STEN	CE P	OSIT	ION:	84.	. 74	9					

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

	GAGO	CGCA	AGG 1	CTGG	GCT	C A	TAGO	TCCC	GGG	CAAC	CGCA	GGC:	rcgc	GC (GGGC	CT	GGG	60
	CGCG	GGA	rcc c	SACTO	TAG	C G1	A A	rg ga	G G	og go	GC GC	C T	TT C	rg ga	AC TO	CG (CTC	113
5							Me	et Gl	.u Al	la Gl	ly G	ly Pl	ne Le	eu As	sp Se	er 1	Leu	
								1				5					10	
	ATT	TAC	GGA	GCA	TGC	GTG	GTC	TTC	ACC	CTT	GGC	ATG	TTC	TCC	GCC	GG	C	161
	Ile	Tyr	Gly	Ala	Ċys	Val	Va1	Phe	Thr	Leu	Gly	Met	Phe	Ser	Ala	Gl:	У	
					15					20					25			
10	CTC	TCG	GAC	CTC	AGG	CAC	ATG	CGA	ATG	ACC	CGG	AGT	GTG	GAC	AAC	GT	2	209
	Leu	Ser	Asp	Leu	Arg	His	Met	Arg	Met	Thr	Arg	Ser	Val	Asp	Asn	Va.	l	
-				30					35					40				
															TGG			257
	Gln	Phe	Leu	Pro	Phe	Leu	Thr	Thr	Glu	Val	Asn	Asn	Leu	Gly	Trp	Le	ı	
15			45					50					55					
															AAC			305
	Ser		Gly	Ala	Leu	Lys		Asp	Gly	Ile	Leu		Val	Vai	Asn	Th:	r	
		60					65				mma	70	m 4 M	C M C	CAT	ጥለ	_	353
00															CAT			223
20		GIÀ	Ala	Ala	Leu		Thr	Leu	Tyr	TTE		ALA	ıyı	reu	His	1 y . 9		
	75 TCC	COM	ccc	4.4.0	CCT	80	CTC	C T C	CTA.	CAC	85	CCA	۸۵۲	ርምር	CTA	-	-	401
															Leu			702
	Cys	110	AL E	Lys	95	Val	Val	Deu	Deu	100	1111	71.44		200	105	,	,	
25	GTC	СТТ	СТС	CTG		ТАТ	GGC	TAC	TTT		CTC	CTG	GTA	ccc	AAC	CC	T	449
															Asn			
				110		-,		-,	115	•				120				
	GAG	GCC	CGG	CTT	CAG	CAG	TTG	GGC	CTC	TTC	TGC	AGT	GTC	TTC	ACC	AT	С	497
	Glu	Ala	Arg	Leu	Gln	Gln	Leu	Gly	Leu	Phe	Cys	Ser	Val	Phe	Thr	11	е	
30			125					130					135					
	AGC	ATG	TAC	CTC	TCA	CCA	CTĢ	GCT	GAC	TTG	GCT	AAG	GTG	ATT	CAA	AC	Т	545
															Gln			
		140					145					150						
	AAA	TCA	ACC	CAA	TGT	CTC	TCC	TAC	CCA	CTC	ACC	ATT	GCT	ACC	CTT	CT	С	593
35	Lys	Ser	Thr	Gln	Cys	Leu	Ser	Tyr	Pro	Leu	Thr	Ile	Ala	Thr	Leu	Le	u	
	155					160					165			-		17	0	
	ACC	TCT	GCC	TCC	TGG	TGC	CTC	TAT	GGG	TTT	CGA	CTC	AGA	GAT	CCC	TA	T	641
	Thr	Ser	Ala	Ser	Trp	Cys	Leu	Tyr	Gly	Phe	Arg	Leu	Arg	Asp	Pro	Ту	r	

	175 180 185	
	ATC ATG GTG TCC AAC TTT CCA GGA ATC GTC ACC AGC TTT ATC CGC TTC	689
	Ile Met Val Ser Asn Phe Pro Gly Ile Val Thr Ser Phe Ile Arg Phe	
	190 195 200	
5	TGG CTT TTC TGG AAG TAC CCC CAG GAG CAA GAC AGG AAC TAC TGG CTC	737
	Trp Leu Phe Trp Lys Tyr Pro Gln Glu Gln Asp Arg Asn Tyr Trp Leu	
	205 210 215	
	CTG CAA ACC TGAGGCTGCT CATCTGACCA CTGGGCACCT TAGTGCCAAC CTGA	790
	Leu Gln Thr	
10	220	
	ACCAAAGAGA CCTCCTTGTT TCAGCTGGGC CTGCTGTCCA GCTTCCCAGG TGCAGTGGGT	850
	TGTGGGAACA AGAGATGACT TTGAGGATAA AAGGACCAAA GAAAAAGCTT TACTTAGATG	910
	ATTGATTGGG GCCTAGGAGA TGAAATCACT TTTTATTTTT TAGAGATTTT TTTTTTTAAT	970
	TTTGGAGGTT GGGGTGCAAT CTTTAGAATA TGCCTTAAAA GGCCGGGCGC GGTGGCTCAC	1030
15	GCCTGTAATC CCAGCACTTT GGGAGGCCAA GGTGGGCGGA TCGCCTGAGG TCAGGAGTTC	1090
	AAGACCAACC TGACTAACAT GGTGAAACCC CATCTCTACT AAAAATACAA AATTAGCCAG	1150
	GCATGATGGC ACATGCCTGT AATCCCAGAT ACTTGGGAGG CTGAGGCAGG AGAATTGCTT	1210
	GAACCCAGGA GGTGGAGGTT GCAGTGAGCT GAGATCGTGC CATTGTGATA TGAATATGCC	1270
	TTATATGCTG ATATGAATAT GCCTTAAAAT AAAGTGTTCC CCACCCCTGC CC	1322
20		
	AGA TARRONALITAN DOR ORO TO NO. AG	
	(2) INFORMATION FOR SEQ ID NO: 42:	
	(i) SEQUENCE CHARACTERISTICS:	
. -	(A) LENGTH: 3045	
25	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear (ii) SEQUENCE KIND: cDNA to mRNA	
	(II) SEQUENCE KIND: CDNA CO HKNA	
30	(vi) ORIGINAL SOURCE:	
-	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Stomach cancer	
	(D) CLONE NAME: HP10230	
	(2) 000112 111210 111 20000	
35	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
	(B) EXISTENCE POSITION: 191 946	
	(C) CHARACTERIZATION METHOD: E	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

	GTTI	rcgcc	CTC A	AGAA	GGCT	GC C	TCGC	TGGT	CCG	AATT	CGGT	GGC	GCCA	CGT	CCGC	CCGTC	т 60
	CCGC	CTTC	CTG (CATC	GCGG	CT T	CGGC	GGCT	T CC	ACCT	AGAC	ACC	TAAC.	AGT (CGCG	GAGCC	G 120
5	GCCG	CGT	CGT (AGG	GGGT	CG G	CACG	GGGA	G TC	GGGC	GGTC	TTG	TGCA	TCT	TGGC	TACCT	G 180
	TGG	TCGA	AAG A	ATG	TCG (GAC .	ATC	GGA	GAC	TGG	TTC	AGG .	AGC .	ATC	CCG	GCG	229
			ì	1et	Ser .	Asp	Ile	Gly	Asp	Trp	Phe	Arg	Ser	Ile :	Pro	Ala	
				1				5					10				
	ATC	ACG	CGC	TAT	TGG	TTC	GCC	GCC	ACC	GTC	GCC	GTG	CCC	TTG	GTC	GGC	277
10	Ile	Thr	Arg	Tyr	Trp	Phe	Ala	Ala	Thr	. Val	Ala	Val	Pro	Leu	Val	Gly	
		15					20	i				25					
	AAA	CTC	GGC	CTC	ATC	AGC	CCG	GCC	TAC	CTC	TTC	CTC	TGG	CCC	GAA	GCC	325
	Lys	Leu	Gly	Leu	Ile	Ser	Pro	Ala	Туг	Leu	ı Phe	Leu	Trp	Pro	Glu	Ala	
	30					35					40)				45	
15	TTC	CTT	TAT	CGC	TTT	CAG	ATT	TGG	AGG	CCA	ATC	ACT	GCC	ACC	TTT	TAT	373
	Phe	Leu	Tyr	Arg	Phe	Gln	Ile	Trp	Arg	g Pro	Ile	Thr	Ala	Thr	Phe	Tyr	
					50					55	j				60	ı	
	TTC	CCT	GTG	GGT	CCA	GGA	ACT	GGA	TTI	CTI	TAT	TTG	GTC	AAT	TTA	TAT	421
	Phe	Pro	Val	Gly	Pro	Gly	Thr	G1y	Phe	e Leu	Туг	Leu	Val	Asn	Leu	Tyr	
20				65					70					75			
	TTC	TTA	TAT	CAG	TAT	TCT	ACG	CGA	CT	GAA	ACA	GGA	GCT	TTT	GAT	GGG	469
	Phe	Leu	Tyr	Gln	Tyr	Ser	Thr	Arg	Lei	ı Glu	Thr	Gly	Ala	Phe	Asp	Gly	
			80					85					90				
																ATC	517
25	Arg	Pro	Ala	Asp	Tyr	Leu	Phe	Met	. Leı	ı Lev	. Phe	Asn	Trp	Ile	Cys	Ile	
		95					100					105	•				
																CTG	565
	Val	Ile	Thr	Gly	Leu	Ala	Met	Asp	Met	Glr	ı Leu	Leu	Met	Ile	Pro	Leu	
	110					115					120					125	
30																ATT	613
	Ile	Met	Ser	Val	Leu	Tyr	Val	Trp	Ala	a Glr	ı Leu	. Asn	Arg	Asp	Met	Ile	
					130					135					140		
																TGG	661
	Val	Ser	Phe	Trp	Phe	Gly	Thr	Arg	Phe	e Lys	: Ala	Cys	Tyr	Leu	Pro	Trp	
35				145					150					155			
																GAG	709
	Val	Ile	Leu	Gly	Phe	Asn	Tyr	Ile	: Ile	e Gly	7 Gly	Ser	Val	Ile	Asn	Glu	
			160					165	,				170				

	CTT	ATT	GGA	AAT	CTG	GTT	GGA	CAT	CTT	TAT	TTT	TTC	CTA	ATG	TTC	AGA	757
	Leu	Ile	Gly	Asn	Leu	Val	Gly	His	Leu	Tyr	Phe	Phe	Leu	Met	Phe	Arg	
		175					180					185					
	TAC	CCA	ATG	GAC	TTG	GGA	GGA	AGA	AAT	TTT	CTA	TCC	ACA	CCT	CAG	TTT	805
5	Tyr	Pro	Met	Asp	Leu	Gly	Gly	Arg	Asn	Phe	Leu	Ser	Thr	Pro	Gln	Phe	
	190					195					200					205	
•	TTG	TAC	CGC	TGG	CTG	ccc	AGT	AGG	AGA	GGA	GGA	GTA	TCA	GGA	TTT	GGT	853
	Leu	Tyr	Arg	Trp	Leu	Pro	Ser	Arg	Arg	Gly	Gly	Val	Ser	Gly	Phe	Gly	
					210					215					220		
10	GTG	ccc	CCT	GCT	AGC	ATG	AGG	CGA	GCT	GCT	GAT	CAG	AAT	GGC	GGA	GGC	901
	Val	Pro	Pro	Ala	Ser	Met	Arg	Arg	Ala	Ala	Asp	Gln	Asn	Gly	Gly	Gly	
				225					230					235			
	GGG	AGA	CAC	AAC	TGG	GGC	CAG	GGC	TTT	CGA	CTT	GGA	GAC	CAG	TGA	AGGG	950
	Gly	Arg	His	Asn	Trp	Gly	Gln	Gly	Phe	Arg	Leu	Gly	Asp	Gln			
15			240					245					250				
																GCTTAA	1010
																CAGTAC	1070
																GATTCT	1130
																CTGACT	1190
20																GGGTCC	1250
																CTTATC	1310
																TAGAAG	1370
•																TGCCAA	1430
																GTAGCA	1490
25																TTCGAC	1550
																CCACTG	1610 1670
																GTTTGT	1730
•																ACCTCC	1790
																AGCTGG	
30																ATGCTC	
																TTCATT	
																CCCCCG	
																TAGATO	
2 -																AATGGC	
35																TCTGTG	
																CAGAGC	
																TTTATT CTTTGA	
	TTA	TGAC	1 - T'T	ATCT	ιτΑΑΑ	GC A	GACT	CTTA	L FLA	ι÷ι.ΑG	IAII	عاΑما	コピピじ	101	UNUA	ヘエエエロな	2000

133

	GGCAACTAAA	AAGGCTTCAA	ACGTTTTGAT	CAGTTTCTTT	TCAGGAAACA	TTGTGCTCTA	2390
	ACAGTATGAC	TATTCTTTCC	CCCACTCTTA	AACAGTGTGA	TGTGTGTTAT	CCTAGGAAAT	2450
	GAGAGTTGGC	AAACAACTTC	TCATTTTGAA	TAGAGTTTGT	GTGTACCTCT	CCATATTTAA	2510
	TTTATATGAT	AAAATAGGTG	GGGAGAGTCT	GAACCTTAAC	TGTCATGTTT	TGTTGTTCAT	2570
5	CTGTGGCCAC	AATAAAGTTT	ACTTGTAAAA	TTTTAGAGGC	CATTACTCCA	ATTATGTTGC	2630
	ACGTACACTC	ATTGTACAGG	CGTGGAGACT	CATTGTATGT	ATAAGAATAT	TCTGACAGTG	2690
	AGTGACCCGG	AGTCTCTGGT	GTACCCTCTT	ACCAGTCAGC	TGCCTGCGAG	CAGTCATTTT	2750
	TTCCTAAAGG	TTTACAAGTA	TTTAGAACTC	TTCAGTTCAG	GGCAAAATGT	TCATGAAGTT	2810
	ATTCCTCTTA	AACATGGTTA	GGAAGCTGAT	GACGTTATTG	ATTTTGTCTG	GATTATGTTT	2870
10	CTGGAATAAT	TTTACCAAAA	CAAGCTATTT	GAGTTTTGAC	TTGACAAGGC	AAAACATGAC	2930
	AGTGGATTCT	CTTTACAAAT	TGAAAAAAA	AATCCTTATT	TTGTATAAAG	GACTTCCCTT	2990
	TTTGTAAACT	AATCCTTTTT	ATTGGTAAAA	ATTGTAAATT	AAAATGTGCA	ACTTG	3045
				_			
15		TION FOR SE					
	(i) S	EQUENCE CHA		CS:			
		(A) LENGTH					
			Nucleic ac				
••			EDNESS: Do	np16			
20		• •	GY: Linear	~ D N A			
	(11)	SEQUENCE KI	.ND: CDNA C	D HENNA			
	(vi)	ORIGINAL SO	URCE:				
	, ,		ISM: Homo s	apiens			
25		(B) CELL E	CIND: Epide:	rmoid carci	noma		
		(C) CELL I					
		(D) CLONE	NAME: HP10	389			
	(ix)	SEQUENCE CH	ARACTERIST:	ics:			
30		(A) CHARAC	CTERIZATION	CODE: CDS			
		(B) EXISTE	ENCE POSITI	ON: 63 38	3		
		(C) CHARAC	CTERIZATION	METHOD: E			
					4.0		
25	(xi)	SEQUENCE DE	ESCRIPTION:	PEG ID MO:	43:		
35	. mo . commc :	00000:000	0.4.00.00000000000000000000000000000000	mcccc + mm===	0m00m00m00	тстсссссс	.60
		ACT CCC GG				TGTGGCCCGG	107
	AC ATTA (40)(4	AUT UGG GG	. L.C.T (+73. A	LI LALAGE LEAGE	いっし しいし エエエ	Our con	±0,

Met Ala Thr Pro Gly Pro Val Ile Pro Glu Val Pro Phe Glu Pro

		1				5					10					15		
	TCG	AAG	CCT	CCA	GTC	ATT	GAG	GGG	CTG	AGC	CCC	ACT	GTT	TAC	AGG	AAT	:	155
	Ser	Lys	Pro	Pro	Val	Ile	Glu	Gly	Leu	Ser	Pro	Thr	Val	Tyr	Arg	Asn		
					20					25					30			
5	CCA	GAG	AGT	TTC	AAG	GAA	AAG	TTC	GTT	CGC	AAG	ACC	CGC	GAG	AAC	CCG	:	203
	Pro	Glu	Ser	Phe	Lys	Glu	Lys	Phe	Val	Arg	Lys	Thr	Arg	Glu	Asn	Pro		
				35					40					45				
															TAC		:	251
	Val	Val	Pro	Ile	Gly	Cys	Leu	Ala	Thr	Ala	Ala	Ala	Leu	Thr	Tyr	Gly		
10			50					55					60					
															ATG			299
	Leu	Tyr	Ser	Phe	His	Arg	Gly	Asn	Ser	Gln	Arg		Gln	Leu	Met	Met		
		65					70					75			mmo	0.00		247
															TTG			347
15	_	Thr	Arg	Ile	Ala		Gln	Gly	Phe	Thr		ALB	ATA	11e	Leu	95		
	80			0.00	4.00	85	4 mo		mcm.	CC 4	90	T A A I	ccc.	۸۵۵	ርጥር ጥ (400
												IAA	3000	noo	G101	GCCTT		400
	Gly	ren	Ala	vaı	100	WIR	met	Lys	261	105	FIU							
20	GAA	ልሮሮሞ∉	CCG	CAGA		ልጥ ጥ	CCAA	AACC	C AG		CAAC	CAC'	TGGC	CCT	ACCG'	TGGGAC		460
20																TTTGTG		520
																CATACT		580
																CACTTG		640
		TTAA																653
25																		
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	44:									
		(i) S	EQUE	NCE	CHAR	ACTE	RIST	ics:									
		•		(A)	LEN	GTH:	439											
30				(B)	TYP	E: N	ucle	ic a	cid									
				(C)	STR	ANDE	DNES	S: D	oubl	е								
				(D)	TOP	OLOG	Y: L	inea	r									
		(ii)	SEQU	ENCE	KIN	D: c	DNA	to m	RNA								•
35		(vi)	ORIG	INAL	sou	RCE:											
				(A)	ORG	ANIS	M: <i>H</i>	iomo	sapi	ens								
				(B)	CEL	L KI	ND:	Stom	ach	canc	er							

(D) CLONE NAME: HP10408

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(ix) SEQUENCE CHARACTERISTICS:

	(A)	CHARACTERIZA	ATION COL	E: CDS			
	(B)	EXISTENCE PO	OSITION:	75 311	L		
	(C)	CHARACTERIZA	ATION MET	HOD: E			
5							
	(xi) SEQU	ENCE DESCRIP	TION: SEC	ID NO:	44:		
	GTAGAAACAG GCCT						60
	GCGCAGTGGC CACT					_	110
10		•	•	Pro Leu	Val Leu	Leu Leu Thr	
		1	5			10	1.50
	CTC CTT GGC AGC						158
	Leu Leu Gly Ser	Ser His Gly		Pro Gly		Leu Gin Leu	
	15		20	maa maa	25	moc 100 mmc	206
15	AAG CTG AAG GAG						200
	Lys Leu Lys Glu		Inr Asn	Ser Ser	40	ser ser rue	
	30 CTG GAA TTG CTT	35	መርር ርሞር	ርምር ርጥር		CCT TCA GGG	254
	Leu Glu Leu Leu						254
20	45	50	cys Leu	55	mis bea	60	
20	ACC AGC GTC ACC		CCA ACA		CAC CAT		302
	Thr Ser Val Thr						• • • • • • • • • • • • • • • • • • • •
	IIII Ser var IIII	65	nra mra	70		75	
	AAC ACA TGACAGC		GT GTCCT1	•	CCGGGCTT		360
25	Asn Thr		0. 0.001.				
23	2.5						
	TGCAGGAGGC AGGC	CCCGAC CCTGT	CTTTC AGO	CAGGCCCC	CACCCTC	TG AGTGGCAATA	420
	AATAAAATTC GGTA						439
30							
	(2) INFORMATION	FOR SEQ ID	NO: 45:				
	(i) SEQUE	NCE CHARACTE	RISTICS:				
	(A)	LENGTH: 113	1				
	(B)	TYPE: Nucle	ic acid				
35	(C)	STRANDEDNES	S: Double	•			
	(D)	TOPOLOGY: L	inear				
	(ii) SEQU	ENCE KIND: c	DNA to m	RNA			

		(v	i) 0	RIGI	NAL	SOUR	CE:										
				(A)	ORGA	NISM	l: <i>Hc</i>	mo s	apie	ns							
				(B)	CELL	KIN	D: S	toma	ch c	ance	r						
				(D)	CLON	E NA	ME:	HP10	412								
5																	
		(i	.x) S	EQUE	NCE	CHAR	ACTE	RIST	ics:								
				(A)	CHAR	ACTE	RIZA	TION	COD	E: C	DS						
				(B)	EXIS	TENC	E PC	SITI	ON:	56	100	0					
				(C)	CHAR	ACTE	RIZA	MOIT.	MET	'HOD:	Ė						
10																	
		()	i) S	EQUE	ENCE	DESC	RIPI	NOI:	SEQ	ID	NO:	45:					
																	50
	CTAI	GAGA	ATC (CCGGC	CCTCA	G GG	TGGA	CGCA	GTG	GTTC	TGC	ACTO	AGGC	CC 1	CGTC	ATG	58
																Met	
15									000	000	000	OBO.	C T A	CTC	CCC	1	106
									GCG								100
	Val	Ala	Pro		Trp	Tyr	Leu	var	Ala 10	AIA	Ala	Leu	ьец	15	019	1	
	4 m.C	CTC.	መጥር	5 CTC	۸.C.T	ccc	٨٥٥	ccc	GGC	ccc	GCG	GCA	TCA		GGC	CAA	154
20									Gly								
20	116	Leu	20	Deu	1111	n. g	501	25	01)	••••			30		,		
	GAG	CCA		CAC	AAT	GAG	GAG		GCA	GGA	GCA	GGC	CGG	GTG	GCC	CAG	202
									Ala								
		35					40					45					
25	CCT	GGG	CCC	CTG	GAG	CCT	GAG	GAG	CCG	AGA	GCT	GGA	GGC	AGG	CCT	CGG	250
	Pro	Gly	Pro	Leu	Glu	Pro	Glu	Glu	Pro	Arg	Ala	Gly	Gly	Arg	Pro	Arg	
	50					55					60					65	
	CGC	CGG	AGG	GAC	CTG	GGC	AGC	CGC	CTA	CAG	GCC	CAG	CGT	CGA	GCC	CAG	298
	Arg	Arg	Arg	Asp	Leu	Gly	Ser	Arg	Leu	Gln	Ala	Gln	Arg	Arg	Ala	Gln	
30					70					75					80		
									GAG								346
	Arg	Val	Ala	Trp	Ala	Glu	Ala	Asp	Glu	Asn	Glu	Glu	Glu	Ala	Val	Ile	
				85					90					95			
									GAG								394
35	Leu	Ala	Gln	Glu	Glu	Glu	Gly	Val	Glu	Lys	Pro	Ala		Thr	His	Leu	
			100					105					110			044	
	TCG	GGG	AAA	ATT					CTG							_	442
	0	C1	*	T1 -	C 1	A 3 -	T	1	7 011	A ** 0	I 37 C	1.011	(i i i i	(÷ [11	1.75	(+ I T)	

		115					120					125					
	GCG	CGA	AAG	GCC	CAG	CGT	GAG	GCA	GAG	GAG	GCT	GAA	CGT	GAG	GAG	CGG	490
	Ala	Arg	Lys	Ala	Gln	Arg	Glu	Ala	Glu	Glu	Ala	Glu	Arg	Glu	Glu	Arg	
	130					135					140					145	
5	AAA	CGA	CTC	GAG	TCC	CAG	CGC	GAA	GCT	GAG	TGG	AAG	AAG	GAG	GAG	GAG	538
	Lys	Arg	Leu	Glu	Ser	Gln	Arg	Glu	Ala	Glu	Trp	Lys	Lys	Glu	Glu	Glu	
					150					155					160		
	CGG	CTT	CGC	CTG	GAG	GAG	GAG	CAG	AAG	GAG	GAG	GAG	GAG	AGG	AAG	GCC	586
	Arg	Leu	Arg	Leu	Glu	Glu	Glu	Gln	Lys	Glu	Glu	Glu	G1u	Arg	Lys	Ala	
10				165					170					175			
	CGC	GAG	GAG	CAG	GCC	CAG	CGG	GAG	CAT	GAG	GAG	TAC	CTG	AAA	CTG	AAG	634
•	Arg	Glu	Glu	Gln	Ala	Gln	Arg	Glu	His	Glu	Glu	Tyr	Leu	Lys	Leu	Lys	
			180					185					190				
	GAG	GCC	TTT	GTG	GTG	GAG	GAG	GAA	GGC	GTA	GGA	GAG	ACC	ATG	ACT	GAG	682
15	Glu	Ala	Phe	Val	Val	Glu	Glu	Glu	Gly	Val	Gly	Glu	Thr	Met	Thr	Glu	
		195					200					205					
								ACA									730
	Glu	Gln	Ser	Gln	Ser	Phe	Leu	Thr	Glu	Phe	Ile	Asn	Tyr	Ile	Lys		
	210					215					220					225	
20								GAC									778
	Ser	Lys	Val	Val	Leu	Leu	Glu	Asp	Leu		Ser	Gln	Val	Gly		Arg	
					230					235			_		240		200
								ATC									826
	Thr	Gln	Asp		Ile	Asn	Arg	Ile		Asp	Leu	Leu	Ala		Gly	Thr	
25				245					250					255		004	074
								CGG									874
	Ile	Thr	-	Val	Ile	Asp	Asp	Arg	Gly	Lys	Phe	Ile		TTe	Tnr	Pro	
			260					265					270	000	000	CMC	022
								AAC									922
30	Glu		Leu	Ala	Ala	Val		Asn	Phe	TTe	Arg		Arg	GIY	Arg	vai	
		275					280		400		шоо	285	4 m.c	000	TICC.	ccc	970
								GCC									370
		TTE	Ala	GIU	Leu		Gin	Ala	ser	ASI		Leu	TIE	MIG	ırþ	305	
2 =	290	C 4 C	# CO		ccc	295	000	004	ccc	TIC A	300	ለ ሮሞ <i>(</i>	ጉር ጥጥ/	ጉ ጉ	-π π ₁		1020
35								CCA		1 GA		noi (JU111	UUU11	١١ د ٠		1020
	AIG	GIU	ser	rro			WIR	Pro	WIH								
	4.00	0.4.0.4.	O III M	00.50	310		0000	0004	m	4 m/n=	un C x un	000	maca.	C A C .	ር ል ጥር ፡	ጉፕርርርር	1080
	ACT	CAGA	GTT I	GG TG	1660	UI A	UUTG	GUTA	AC.	MICT	TOWL	CCC	1000	CAC (CAIO	CTGGGG	1000

PCT/JP98/02445 WO 98/55508

303

138

	AAGTGATGGT GTGGCCAGGC AGTTATAGAT TAAAGGCCTG TGAGTACTGC T	1131												
	(2) INFORMATION FOR SEQ ID NO: 46:													
5	(i) SEQUENCE CHARACTERISTICS:													
	(A) LENGTH: 1875													
	(B) TYPE: Nucleic acid													
	(C) STRANDEDNESS: Double													
	(D) TOPOLOGY: Linear													
10	(ii) SEQUENCE KIND: cDNA to mRNA													
	(vi) ORIGINAL SOURCE:													
	(A) ORGANISM: Homo sapiens													
	(B) CELL KIND: Stomach cancer													
15	(D) CLONE NAME: HP10413													
	(ix) SEQUENCE CHARACTERISTICS:													
	(A) CHARACTERIZATION CODE: CDS													
	(B) EXISTENCE POSITION: 79 666													
20	(C) CHARACTERIZATION METHOD: E													
	·													
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:													
	CTCGCTCGCT CAGAGGGAGG AGAAAGTGGC GAGTTCCGGA TCCCTGCCTA GCGCGGCCCA	60												
25	ACCTTTACTC CAGAGATC ATG GCT GCC GAG GAT GTG GTG GCG ACT GGC GCC	111												
	Met Ala Ala Glu Asp Val Val Ala Thr Gly Ala													
	1 5 10													
	GAC CCA AGC GAT CTG GAG AGC GGC GGG CTG CTG CAT GAG ATT TTC ACG	159												
	Asp Pro Ser Asp Leu Glu Ser Gly Gly Leu Leu His Glu Ile Phe Thr													
30	15 20 25													
	TCG CCG CTC AAC CTG CTG CTT GGC CTC TGC ATC TTC CTG CTC TAC	207												
	Ser Pro Leu Asn Leu Leu Leu Gly Leu Cys Ile Phe Leu Leu Tyr													
	30 35 40													
	AAG ATC GTG CGC GGG GAC CAG CCG GCC GCC GGC GAC AGC GAC GA	255												
35	Lys Ile Val Arg Gly Asp Gln Pro Ala Ala Ser Gly Asp Ser Asp Asp													
	45 50 55													

GAC GAG CCG CCC CTG CCC CGC CTC AAG CGG CGC GAC TTC ACC CCC

Asp Glu Pro Pro Pro Leu Pro Arg Leu Lys Arg Arg Asp Phe Thr Pro

	60					65					70					13	
	GCC	GAG	CTG	CGG	CGC	TTC	GAC	GGC	GTC	CAG	GAC	CCG	CGC	ATA	CTC	ATG	351
	Ala	Glu	Leu	Arg	Arg	Phe	Asp	Gly	Val	Gln	Asp	Pro	Arg	Ile	Leu	Met	
					80					85					90		
5	GCC	ATC	AAC	GGC	AAG	GTG	TTC	GAT	GTG	ACC	AAA	GGC	CGC	AAA	TTC	TAC	399
	Ala	Ile	Asn	Gly	Lys	Val	Phe	Asp	Val	Thr	Lys	Gly	Arg	Lys	Phe	Tyr	
				95					100					105			
	GGG	ccc	GAG	GGG	CCG	TAT	GGG	GTC	TTT	GCT	GGA	AGA	GAT	GCA	TCC	AGG	447
	Gly	Pro	Glu	Gly	Pro	Tyr	Gly	Val	Phe	Ala	Gly	Arg	Asp	Ala	Ser	Arg	
LO			110					115					120				
	GGC	CTT	GCC	ACA	TTT	TGC	CTG	GAT	AAG	GAA	GCA	CTG	AAG	GAT	GAG	TAC	495
	Gly	Leu	Ala	Thr	Phe	Cys	Leu	Asp	Lys	Glu	Ala	Leu	Lys	Asp	G1u	Tyr	
		125					130					135					
	GAT	GAC	CTT	TCT	GAC	CTC	ACT	GCT	GCC	CAG	CAG	GAG	ACT	CTG	AGT	GAC	543
15	Asp	Asp	Leu	Ser	Asp	Leu	Thr	Ala	Ala	Gln	Gln	Glu	Thr	Leu	Ser	Asp	
	140					145					150					155	
	TGG	GAG	TCT	CAG	TTC	ACT	TTC	AAG	TAT	CAT	CAC	GTG	GGC	AAA	CTG	CTG	591
	Trp	Glu	Ser	Gln	Phe	Thr	Phe	Lys	Tyr	His	His	Val	Gly	Lys	Leu	Leu	
					160					165					170		
20	AAG	GAG	GGG	GAG	GAG	ccc	ACT	GTG	TAC	TCA	GAT	GAG	GAA	GAA	CCA	AAA	639
	Lys	Glu	Gly	Glu	Glu	Pro	Thr	Val	Tyr	Ser	Asp	Glu	Glu	Glu	Pro	Lys	
				175					180					185			
	GAT	GAG	AGT	GCC	CGG	AAA	TAA	GAT	TAA	AGCA	TTC A	AGTG(GAAG'	ra T.	ATCT	AT	690
	Asp	Glu	Ser	Ala	Arg	Lys	Asn	Asp									
25			190					195									
																GATTAC	750
																ACACTA	810
																GTGAAC	870
																CAGGAT	930
30																CAGGTA	990
																GTAAAG	1050
																TCTACC	1110
																AGTAAC	1170
																CTGGAC	1230
35																AGTCTA	1290
																AGCTGG	1350
	CTG	GCCT	AAA	AACC	TAAA	TA T	'ATGA	TGAA	G AT	TGTA	GGAC	TGT	CTTC	CCA	AGCC	CCATGT	1410
	Tr.C.A	ጥርርጥ	ccc	CCAA	TOOT	ጥ ለ ጥ	ישייכב	ጥጥልጥ	ጥ ጥጥ	АСТС	ΑΑΤΤ	CCT	TACT	CTC	ATTT	GAAATG	1470

140

AGGGAGGGAC ATACAGAATA GGAACAGGTG TTTGCTCTCC TAAGAGCCTT CATGCACACC 1530

	CCTGAACCAC GAGGAAACAG TACAGTCGCT AGTCAAGTGG TTTTTAAAGT AAAGTATATT	1590
	CATAAGGTAA CAGTTATTCT GTTGTTATAA AACTATACCC ACTGCAAAAG TAGTAGTCAA	1650
	GTGTCTAGGT CTTTGATATT GCTCTTTTGG TTAACACTAA GCTTAAGTAG ACTATACAGT	1710
5	TGTATGAATT TGTAAAAGTA TATGAACACC TAGTGAGATT TCAAACTTGT AATTGTGGTT	1770
	AAATAGTCAT TGTATTTTCT TGTGAACTGT GTTTTATGAT TTTACCTCAA ATCAGAAAAC	1830
	AAAATGATGT GCTTTGGTCA GTTAATAAAA ATGGTTTTAC CCACT	1875
10	(2) INFORMATION FOR SEQ ID NO: 47:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1563	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
15	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(VI) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens	
20	(B) CELL KIND: Stomach cancer	
. 20	(D) CLONE NAME: HP10415	
	(2, 52002 00020 00000	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
25	(B) EXISTENCE POSITION: 72 1460	
	(C) CHARACTERIZATION METHOD: E	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:	
		60
30	AAATTGGGCC AGGCTGAGGC GCTGCTGCTG GAGCGGCCGA TCCGAGACGT GGCTCCCTGG	60
	GCGGCAGAAC C ATG TTG GAC TTC GCG ATC TTC GCC GTT ACC TTC TTG CTG	110
	Met Leu Asp Phe Ala Ile Phe Ala Val Thr Phe Leu Leu	
	1 5 10	158
2 =	GCG TTG GTG GGA GCC GTG CTC TAC CTC TAT CCG GCT TCC AGA CAA GCT	250
35	Ala Leu Val Gly Ala Val Leu Tyr Leu Tyr Pro Ala Ser Arg Gln Ala	
	GCA GGA ATT CCA GGG ATT ACT CCA ACT GAA GAA AAA GAT GGT AAT CTT	206
	Ale Civ Tie Pro Civ Tie Thr Pro Thr Glu Glu Lys Asp Gly Asn Leu	.= - -

	30					35					40					45	
	CCA	GAT	ATT	GTG	TAA	AGT	GGA	AGT	TTG	CAT	GAG	TTC	CTG	GTT	TAA	TTG	254
	Pro	Asp	Ile	Val	Asn	Ser	Gly	Ser	Leu	His	Glu	Phe	Leu	Val	Asn	Leu	
					50					55					60		
5	CAT	GAG	AGA	TAT	GGG	CCT	GTG	GTC	TCC	TTC	TGG	TTT	GGC	AGG	CGC	CTC	302
	His	Glu	Arg	Tyr	Gly	Pro	Val	Val	Ser	Phe	Trp	Phe	Gly	Arg	Arg	Leu	
				65					70					75			
	GTG	GTT	AGT	TTG	GGC	ACT	GTT	GAT	GTA	CTG	AAG	CAG	CAT	ATC	AAT	CCC	350
	Val	Val	Ser	Leu	Gly	Thr	Val	Asp	Val	Leu	Lys	Gln	His	Ile	Asn	Pro	
10			80					85					90				
	AAT	AAG	ACA	TTG	GAC	CCT	TTT	GAA	ACC	ATG	CTG	AAG	TCA	TTA	TTA	AGG	398
•	Asn	Lys	Thr	Leu	Asp	Pro	Phe	Glu	Thr	Met	Leu	Lys	Ser	Leu	Leu	Arg	
		95					100					105					
									AGT								446
15	Tyr	Gln	Ser	Gly	Gly	Gly	Ser	Val	Ser	Glu	Asn	His	Met	Arg	Lys		
	110					115					120					125	
									TCT								494
	Leu	Tyr	Glu	Asn		Val	Thr	Asp	Ser		Lys	Ser	Asn	Phe		Leu	
					130					135				maa	140	664	5
20									TTA								542
	Leu	Leu	Lys		Ser	Glu	Glu	Leu	Leu	Asp	Lys	Trp	Leu		Tyr	PIO	
				145					150	O 4 M	4.TO	C TI TI	CC#	155	CCT	ΔTG	590
									CAG								330
25	GIU	Thr			val	Pro	Leu	165	Gln	urs	Met	Leu	170	THE	niu	1100	
25	440	ጥርጥ	160		CAC	ልጥር	GTΔ		GGT	ΔСТ	ACA	ጥጥጥ		GAT	GAT	CAG	638
									Gly								
	Lys	175		1111	GIII	nec	180	nec	019	501	****	185	014				
	GAA			CGC	TTC	CAG		ΑΑΤ	CAT	GGC	ACA		TGG	TCT	GAG	ATT	686
30									His								
30	190					195					200		•			205	
		AAA	GGC	TTT	CTA			TCA	CTT	GAT	AAA	AAC	ATG	ACT	CGG	AAA	734
									Leu								
	•	•	•		210		·			215					220		
35	AAA	CAA	TAT	GAA	GAT	GCC	CTC	ATG	CAA	CTG	GAG	TCT	GTT	TTA	AGG	AAC	782
									Gln								
	-		-	225					230					235			
	ATC	ATA	AAA	GAA	CGA	AAA	GGA	AGG	AAC	TTC	AGT	CAA	CAT	ATT	TTC	ATT	830

	Ile	Ile	Lys	G1u	Arg	Lys	Gly	Arg	Asn	Phe	Ser	Gln	His	Ile	Phe	Ile	
			240					245					250				
	GAC	TCC	TTA	GTA	CAA	GGG	AAC	CTT	TAA	GAC	CAA	CAG	ATC	CTA	GAA	GAC	878
	Asp	Ser	Leu	Val	Gln	Gly	Asn	Leu	Asn	Asp	Gln	Gln	Ile	Leu	Glu	Asp	
5		255					260					265					
	AGT	ATG	ATA	TTT	TCT	CTG	GCC	AGT	TGC	ATA	ATA	ACT	GCA	AAA	TTG	TGT	926
	Ser	Met	Ile	Phe	Ser	Leu	Ala	Ser	Cys	Ile	Ile	Thr	Ala	Lys	Leu	Cys	
	270					275					280					285	
	ACC	TGG	GCA	ATC	TGT	TTT	TTA	ACC	ACC	TCT	GAA	GAA	GTT	CAA	AAA	AAA	974
10	Thr	Trp	Ala	Ile	Cys	Phe	Leu	Thr	Thr	Ser	Glu	Glu	Val	Gln	Lys	Lys	
					290					295					300		
	TTA	TAT	GAA	GAG	ATA	AAC	CAA	GTT	TTT	GGA	AAT	GGT	CCT	GTT	ACT	CCA	1022
	Leu	Tyr	Glu	G1u	Ile	Asn	Gln	Val	Phe	Gly	Asn	Gly	Pro	Val	Thr	Pro	
				305					310					315			
15	GAG	AAA	ATT	GAG	CAG	CTC	AGA	TAT	TGT	CAG	CAT	GTG	CTT	TGT	GAA	ACT	1070
	Glu	Lys	Ile	Glu	Gln	Leu	Arg	Tyr	Cys	Gln	His	Val	Leu	Cys	Glu	Thr	
			320					325					330				
	GTT	CGA	ACT	GCC	AAA	CTG	ACT	CCA	GTT	TCT	GCC	CAG	CTT	CAA	GAT	ATT	1118
	Val	Arg	Thr	Ala	Lys	Leu	Thr	Pro	Val	Ser	Ala	Gln	Leu	Gln	Asp	Ile	
20		335					340					345					
• •	GAA	GGA	AAA	ATT	GAC	CGA	TTT	ATT	ATT	CCT	AGA	GAG	ACC	CTC	GTC	CTT	1166
	Glu	Gly	Lys	Ile	Asp	Arg	Phe	Ile	Ile	Pro	Arg	Glu	Thr	Leu	Val	Leu	
	350					355					360					365	
	TAT	GCC	CTT	GGT	GTG	GTA	CTT	CAG	GAT	CCT	AAT	ACT	TGG	CCA	TCT	CCA	1214
25	Tyr	Ala	Leu	Gly	Val	Val	Leu	Gln	Asp	Pro	Asn	Thr	Trp	Pro	Ser	Pro	
					370					375					380		
•									GAT								1262
	His	Lys	Phe	Asp	Pro	Asp	Arg	Phe	Asp	Asp	Glu	Leu	Val		Lys	Thr	
				385					390					395			
30									ACA								1310
	Phe	Ser		Leu	Gly	Phe	Ser		Thr	Gln	Glu	Cys		Glu	Leu	Arg	
			400					405					410				
									CTT								1358
	Phe	Ala	Tyr	Met	Val	Thr	Thr	Val	Leu	Leu	Ser		Leu	Val	Lys	Arg	
35		415					420					425			m·-	244	3.4.0.0
									CAG								1406
	Leu	His	Leu	Leu	Ser	Val	Glu	Gly	Gln	Val		Glu	Thr	Lys	Tyr		
	430					435					440					445	

	CTG GTA ACA TCA TCA AGG GAA GAA GUT TGG ATU ACT GTO TCA AAG AGA	1424
	Leu Val Thr Ser Ser Arg Glu Glu Ala Trp Ile Thr Val Ser Lys Arg	
	450 455 460	
	TAT TAAAATTTTA TACATTTAAA ATCATTGTTA AATTGATTGA GGAAAACAAC CAT	1510
5	Tyr	
	TTAAAAAAA TCTATGTTGA ATCCTTTTAT AAACCAGTAT CACTTTGTAA TAT	1563
10	(2) INFORMATION FOR SEQ ID NO: 48:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2030	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
15	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	• •	
20	(A) ORGANISM: Homo sapiens (B) CELL KIND: Stomach cancer	
20	(D) CLONE NAME: HP10419	
	(b) Chorn Main. In 19415	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
25	(B) EXISTENCE POSITION: 171 914	
	(C) CHARACTERIZATION METHOD: E	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:	
30	CATTTGGGGT TTCGGTTCCC CCCCTTCCCC TTCCCCGGGG TCTGGGGGTG ACATTGCACC	60
	GCGCCCCTCG TGGGGTCGCG TTGCCACCCC ACGCGGACTC CCCAGCTGGC GCGCCCCTCC	120
	CATTTGCCTG TCCTGGTCAG GCCCCCACCC CCCTTCCCAC CTGACCAGCC ATG GGG	176
	Met Gly	
	1	
35	GCT GCG GTG TTT TTC GGC TGC ACT TTC GTC GCG TTC GGC CCG GCC TTC	224
	Ala Ala Val Phe Phe Gly Cys Thr Phe Val Ala Phe Gly Pro Ala Phe	
	5 10 15	
	GCG CTT TTC TTG ATC ACT GTG GCT GGG GAC CCG CTT CGC GTT ATC ATC	272

	Ala		Phe	Leu	Ile	Thr		Ala	Gly	Asp	Pro		Arg	Val	Ile	Ile	
		20					25					30		0.00		mom	200
					GCA												320
_		Val	Ala	Gly	Ala		Phe	Trp	Leu	val		Leu	Leu	Leu	WIR		
5	35					40					45	000	mc 4	O 4 TT	000	50	260
					ATC												368
	Val	Val	Trp	Phe	Ile	Leu	Val	His	Val		Asp	Arg	Ser	Asp		Arg	
					55					60					65	0.004	126
					CTC												416
10	Leu	Gln	Tyr		Leu	Leu	Ile	Phe		Ala	Ala	Val	Ser		Leu	Leu	
				70					75					80		0.4 m	
					CGC												464
	Gln	Glu		Phe	Arg	Phe	Ala		Tyr	Lys	Leu	Leu		Lys	Ala	Asp	
			85					90					95				
15					TCG												512
	Glu	Gly	Leu	Ala	Ser	Leu	Ser	Glu	Asp	Gly	Arg		Pro	Ile	Ser	116	
		100					105					110					5.60
					TAT												560
	Arg	Gln	Met	Ala	Tyr	Val	Ser	Gly	Leu	Ser	Phe	Gly	Ile	Ile	Ser	Gly	
20	115					120					125					130	
20	GTC				ATC	AAT					GCA					GTG	608
20	GTC				Ile	AAT				Asp	GCA				Gly	GTG	608
20	GTC Val	Phe	Ser	Val	Ile 135	AAT Asn	Ile	Leu	Ala	Asp 140	GCA Ala	Leu	Gly	Pro	Gly 145	GTG Val	
	GTC Val	Phe GGG	Ser ATC	Val CAT	Ile 135 GGA	AAT Asn GAC	Ile	Leu	Ala TAT	Asp 140 TAC	GCA Ala TTC	Leu CTG	Gly ACT	Pro TCA	Gly 145 GCC	GTG Val TTT	608
20	GTC Val	Phe GGG	Ser ATC	Val CAT His	Ile 135	AAT Asn GAC	Ile	Leu	Ala TAT Tyr	Asp 140 TAC	GCA Ala TTC	Leu CTG	Gly ACT	Pro TCA Ser	Gly 145 GCC	GTG Val TTT	
	GTC Val GTT Val	Phe GGG Gly	Ser ATC Ile	Val CAT His 150	Ile 135 GGA Gly	AAT Asn GAC Asp	Ile TCA Ser	Leu CCC Pro	Ala TAT Tyr 155	Asp 140 TAC Tyr	GCA Ala TTC Phe	Leu CTG Leu	Gly ACT Thr	TCA Ser 160	Gly 145 GCC Ala	GTG Val TTT Phe	656
	GTC Val GTT Val	Phe GGG Gly ACA	Ser ATC Ile GCA	Val CAT His 150 GCC	Ile 135 GGA Gly	AAT Asn GAC Asp	TCA Ser	CCC Pro	Ala TAT Tyr 155 CAT	Asp 140 TAC Tyr	GCA Ala TTC Phe	Leu CTG Leu TGG	Gly ACT Thr	TCA Ser 160 GTT	Gly 145 GCC Ala GTG	GTG Val TTT Phe	
	GTC Val GTT Val	Phe GGG Gly ACA	Ser ATC Ile GCA Ala	Val CAT His 150 GCC	Ile 135 GGA Gly	AAT Asn GAC Asp	TCA Ser	Leu CCC Pro CTC Leu	Ala TAT Tyr 155 CAT	Asp 140 TAC Tyr	GCA Ala TTC Phe	Leu CTG Leu TGG	Gly ACT Thr GGA Gly	TCA Ser 160 GTT	Gly 145 GCC Ala GTG	GTG Val TTT Phe	656
25	GTC Val GTT Val CTG Leu	Phe GGG Gly ACA Thr	Ser ATC Ile GCA Ala 165	CAT His 150 GCC Ala	Ile 135 GGA Gly ATT Ile	AAT Asn GAC Asp ATC	TCA Ser CTG Leu	CCC Pro CTC Leu 170	Ala TAT Tyr 155 CAT His	Asp 140 TAC Tyr ACC	GCA Ala TTC Phe TTT Phe	Leu CTG Leu TGG Trp	ACT Thr GGA Gly 175	TCA Ser 160 GTT Val	Gly 145 GCC Ala GTG Val	GTG Val TTT Phe TTC Phe	704
	GTC Val GTT Val CTG Leu	Phe GGG Gly ACA Thr	ATC Ile GCA Ala 165 GCC	CAT His 150 GCC Ala	Ile 135 GGA Gly ATT Ile	AAT ASD ATC Ile	TCA Ser CTG Leu	CCC Pro CTC Leu 170 CGG	Ala TAT Tyr 155 CAT His	Asp 140 TAC Tyr ACC Thr	GCA Ala TTC Phe TTT Phe GCT	Leu CTG Leu TGG Trp	Gly ACT Thr GGA Gly 175 GGC	TCA Ser 160 GTT Val	Gly 145 GCC Ala GTG Val	GTG Val TTT Phe TTC Phe	656
25	GTC Val GTT Val CTG Leu	Phe GGG Gly ACA Thr GAT Asp	ATC Ile GCA Ala 165 GCC	CAT His 150 GCC Ala	Ile 135 GGA Gly ATT Ile	AAT ASD ATC Ile	TCA Ser CTG Leu AGA	CCC Pro CTC Leu 170 CGG	Ala TAT Tyr 155 CAT His	Asp 140 TAC Tyr ACC Thr	GCA Ala TTC Phe TTT Phe GCT	Leu CTG Leu TGG Trp TTG Leu	Gly ACT Thr GGA Gly 175 GGC	TCA Ser 160 GTT Val	Gly 145 GCC Ala GTG Val	GTG Val TTT Phe TTC Phe	704
25	GTC Val GTT Val CTG Leu TTT Phe	Phe GGG Gly ACA Thr GAT Asp 180	ATC Ile GCA Ala 165 GCC Ala	CAT His 150 GCC Ala TGT Cys	Ile 135 GGA Gly ATT Ile GAG Glu	AAT Asn GAC Asp ATC Ile AGG Arg	TCA Ser CTG Leu AGA Arg 185	CCC Pro CTC Leu 170 CGG Arg	Ala TAT Tyr 155 CAT His TAC	Asp 140 TAC Tyr ACC Thr	GCA Ala TTC Phe TTT Phe GCT Ala	Leu CTG Leu TGG Trp TTG Leu 190	Gly ACT Thr GGA Gly 175 GGC Gly	TCA Ser 160 GTT Val CTG Leu	Gly 145 GCC Ala GTG Val	GTG Val TTT Phe TTC Phe GTT Val	704 752
25	GTC Val GTT Val CTG Leu TTT Phe	Phe GGG Gly ACA Thr GAT Asp 180 AGT	Ser ATC Ile GCA Ala 165 GCC Ala CAC	CAT His 150 GCC Ala TGT Cys	Ile 135 GGA Gly ATT Ile GAG Glu CTG	AAT Asn GAC Asp ATC Ile AGG Arg	TCA Ser CTG Leu AGA Arg 185 TCG	CCC Pro CTC Leu 170 CGG Arg	Ala TAT Tyr 155 CAT His TAC Tyr	Asp 140 TAC Tyr ACC Thr TGG Trp	GCA Ala TTC Phe TTT Phe GCT Ala	CTG Leu TGG Trp TTG Leu 190 CTG	Gly ACT Thr GGA Gly 175 GGC Gly AAC	TCA Ser 160 GTT Val CTG Leu CCC	Gly 145 GCC Ala GTG Val GTG Val	GTG Val TTT Phe TTC Phe GTT Val	704
30	GTC Val GTT Val CTG Leu TTT Phe GGG Gly	Phe GGG Gly ACA Thr GAT Asp 180 AGT	Ser ATC Ile GCA Ala 165 GCC Ala CAC	CAT His 150 GCC Ala TGT Cys	Ile 135 GGA Gly ATT Ile GAG Glu	AAT Asn GAC Asp ATC Ile AGG Arg ACA Thr	TCA Ser CTG Leu AGA Arg 185 TCG	CCC Pro CTC Leu 170 CGG Arg	Ala TAT Tyr 155 CAT His TAC Tyr	Asp 140 TAC Tyr ACC Thr TGG Trp	GCA Ala TTC Phe TTT Phe GCT Ala TTC Phe	CTG Leu TGG Trp TTG Leu 190 CTG	Gly ACT Thr GGA Gly 175 GGC Gly AAC	TCA Ser 160 GTT Val CTG Leu CCC	Gly 145 GCC Ala GTG Val GTG Val	GTG Val TTT Phe TTC Phe GTT Val TAT Tyr	704 752
25	GTC Val GTT Val CTG Leu TTT Phe GGG Gly 195	Phe GGG Gly ACA Thr GAT Asp 180 AGT Ser	Ser ATC Ile GCA Ala 165 GCC Ala CAC	CAT His 150 GCC Ala TGT Cys CTA Leu	Ile 135 GGA Gly ATT Ile GAG Glu CTG Leu	AAT Asn GAC Asp ATC Ile AGG Arg ACA Thr 200	TCA Ser CTG Leu AGA Arg 185 TCG Ser	CCC Pro CTC Leu 170 CGG Arg	Ala TAT Tyr 155 CAT His TAC Tyr CTG Leu	Asp 140 TAC Tyr ACC Thr TGG Trp	GCA Ala TTC Phe TTT Phe GCT Ala TTC Phe 205	Leu CTG Leu TGG Trp TTG Leu 190 CTG Leu	Gly ACT Thr GGA Gly 175 GGC Gly AAC Asn	TCA Ser 160 GTT Val CTG Leu CCC	Gly 145 GCC Ala GTG Val GTG TGG	GTG Val TTT Phe TTC Phe GTT Val TAT Tyr 210	704 752
30	GTC Val GTT Val CTG Leu TTT Phe GGG Gly 195 GAG	Phe GGG Gly ACA Thr GAT ASP 180 AGT Ser	Ser ATC Ile GCA Ala 165 GCC Ala CAC His	CAT His 150 GCC Ala TGT Cys CTA Leu CTG	Ile 135 GGA Gly ATT Ile GAG Glu CTG Leu	AAT Asn GAC Asp ATC Ile AGG Arg ACA Thr 200 CCC	TCA Ser CTG Leu AGA Arg 185 TCG Ser	CCC Pro CTC Leu 170 CGG Arg GGA Gly TAT	Ala TAT Tyr 155 CAT His TAC Tyr CTG Leu GCA	Asp 140 TAC Tyr ACC Thr TGG Trp ACA Thr	GCA Ala TTC Phe TTT Phe GCT Ala TTC Phe 205 ACT	CTG Leu TGG Trp TTG Leu 190 CTG Leu	Gly ACT Thr GGA Gly 175 GGC Gly AAC Asn TCC	TCA Ser 160 GTT Val CTG Leu CCC Pro	Gly 145 GCC Ala GTG Val GTG TGG Trp	GTG Val TTT Phe TTC Phe GTT Val TAT Tyr 210 CTC	704 752
30	GTC Val GTT Val CTG Leu TTT Phe GGG Gly 195 GAG	Phe GGG Gly ACA Thr GAT ASP 180 AGT Ser	Ser ATC Ile GCA Ala 165 GCC Ala CAC His	CAT His 150 GCC Ala TGT Cys CTA Leu CTG	Ile 135 GGA Gly ATT Ile GAG Glu CTG Leu	AAT Asn GAC Asp ATC Ile AGG Arg ACA Thr 200 CCC	TCA Ser CTG Leu AGA Arg 185 TCG Ser	CCC Pro CTC Leu 170 CGG Arg GGA Gly TAT	Ala TAT Tyr 155 CAT His TAC Tyr CTG Leu GCA	Asp 140 TAC Tyr ACC Thr TGG Trp ACA Thr	GCA Ala TTC Phe TTT Phe GCT Ala TTC Phe 205 ACT	CTG Leu TGG Trp TTG Leu 190 CTG Leu	Gly ACT Thr GGA Gly 175 GGC Gly AAC Asn TCC	TCA Ser 160 GTT Val CTG Leu CCC Pro	Gly 145 GCC Ala GTG Val GTG TGG Trp	GTG Val TTT Phe TTC Phe GTT Val TAT Tyr 210 CTC	704 752

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	TGG GCC TTC ATC ACA GCT GGA GGG TCC CTC CGA AGT ATT CAG CGC AGC	896
	Trp Ala Phe Ile Thr Ala Gly Gly Ser Leu Arg Ser Ile Gln Arg Ser	
	230 235 240	
	CTC TTG TGT AAG GAC TGACTACCTG GACTGATCGC CTGACAGATC CCACCTGCC	950
5	Leu Leu Cys Lys Asp	
	245	
	TGTCCACTGC CCATGACTGA GCCCAGCCCC AGCCCGGGTC CATTGCCCAC ATTCTCTGTC	1010
	TCCTTCTCGT CGGTCTACCC CACTACCTCC AGGGTTTTGC TTTGTCCTTT TGTGACCGTT	1070
	AGTCTCTAAG CTTTACCAGG AGCAGCCTGG GTTCAGCCAG TCAGTGACTG GTGGGTTTGA	1130
١0	ATCTGCACTT ATCCCCACCA CCTGGGGACC CCCTTGTTGT GTCCAGGACT CCCCCTGTGT	1190
	CAGTGCTCTG CTCTCACCCT GCCCAAGACT CACCTCCCTT CCCCTCTGCA GGCCGACGGC	1250
	AGGAGGACAG TCGGGTGATG GTGTATTCTG CCCTGCGCAT CCCACCCGAG GACTGAGGGA	1310
	ACCTAGGGGG GACCCCTGGG CCTGGGGTGC CCTCCTGATG TCCTCGCCCT GTATTTCTCC	1370
	ATCTCCAGTT CTGGACAGTG CAGGTTGCCA AGAAAAGGGA CCTAGTTTAG CCATTGCCCT	1430
1.5	GGAGATGAAA TTAATGGAGG CTCAAGGATA GATGAGCTCT GAGTTTCTCA GTACTCCCTC	1490
	AAGACTGGAC ATCTTGGTCT TTTTCTCAGG CCTGAGGGGG AACCATTTTT GGTGTGATAA	1550
	ATACCCTAAA CTGCCTTTTT TTCTTTTTTG AGGTGGGGGG AGGGAGGAGG TATATTGGAA	1610
	CTCTTCTAAC CTCCTTGGGC TATATTTTCT CTCCTCGAGT TGCTCCTCAT GGCTGGGCTC	1670
	ATTTCGGTCC CTTTCTCCTT GGTCCCAGAC CTTGGGGGAA AGGAAGGAAG TGCATGTTTG	1730
20	GGAACTGGCA TTACTGGAAC TAATGGTTTT AACCTCCTTA ACCACCAGCA TCCCTCCTCT	1790
	CCCCAAGGTG AAGTGGAGGG TGCTGTGGTG AGCTGGCCAC TCCAGAGCTG CAGTGCCACT	1850
	GGAGGAGTCA GACTACCATG ACATCGTAGG GAAGGAGGGG AGATTTTTTT GTAGTTTTTA	1910
	ATTGGGGTGT GGGAGGGCG GGGAGGTTTT CTATAAACTG TATCATTTTC TGCTGAGGGT	1970
	GGAGTGTCCC ATCCTTTTAA TCAAGGTGAT TGTGATTTTG ACTAATAAAA AAGAATTTGT	2030
25		
		•
	(2) INFORMATION FOR SEQ ID NO: 49:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 493	

- (B) TYPE: Nucleic acid
 - (C) STRANDEDNESS: Double
 - (D) TOPOLOGY: Linear
 - (ii) SEQUENCE KIND: cDNA to mRNA
- 35 (vi) ORIGINAL SOURCE:

- (A) ORGANISM: Homo sapiens
- (B) CELL KIND: Stomach cancer
- (D) CLONE NAME: HP10424

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(A) CHARACTERIZATION CODE: CDS

(ix) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2044

				(B)	EXIS	STEN	E PO	SITI	CON:	98.	439	}					
(C) CHARACTERIZATION METHOD: E																	
5																	
		()	(i)	SEQUE	ENCE	DESC	RIPT	ION:	SEC	Q ID	NO:	49:					
	AAAC	TTTC	CCC A	AAATO	CCAG	GC GC	CTAC	AGG	CCA	ACTG	CTTC	CCA	ACTAC	CCA (GCTG	AGGGGG	60
	TCCC	TCC	CGA (GAAGO	GAG	AA GA	AGGCC	GAAG	AG(GAAA	CATO	AA(TTO	TAT	TT?	A CTC	115
LO																ı Leu	
											1	l				5	
	CTA	GCG	AGC	AGC	ATT	CTG	TGT	GCC	TTG	ATT	GTC	TTC	TGG	AAA	TAT	CGC	163
	Leu	Ala	Ser	Ser	Ile	Leu	Cys	Ala	Leu	Ile	Val	Phe	Trp	Lys	Tyr	Arg	
				10					15					20			
15	CGC	TTT	CAG	AGA	AAC	ACT	GGC	GAA	ATG	TCA	TCA	AAT	TCA	ACT	GCT	CTT	211
	Arg	Phe	Gln	Arg	Asn	Thr	Gly	Glu	Met	Ser	Ser	Asn	Ser	Thr	Ala	Leu	
			25					30					35				
	GCA	CTA	GTG	AGA	ccc	TCT	TCT	TCT	GGG	TTA	ATT	AAC	AGC	AAT	ACA	GAC	259
	Ala	Leu	Val	Arg	Pro	Ser	Ser	Ser	Gly	Leu	Ile	Asn	Ser	Asn	Thr	Asp	
20		40					45					50					
	AAC	AAT	CTT	GCA	GTC	TAC	GAC	CTC	TCT	CGG	GAT	ATT	TTA	AAT	AAT	TTC	307
	Asn	Asn	Leu	Ala	Val	Tyr	Asp	Leu	Ser	Arg	Asp	Ile	Leu	Asn	Asn	Phe	
	55					60					65					70	
	CCA	CAC	TCA	ATA	GCC	AGG	CAG	AAG	CGA	ATA	TTG	GTA	AAC	CTC	AGT	ATG	355
25	Pro	His	Ser	Ile	Ala	Arg	G1n	Lys	Arg	Ile	Leu	Val	Asn	Leu	Ser	Met	
					75					80					85		
	GTG	GAA	AAC	AAG	CTG	GTT	GAA	CTG	GAA	CAT	ACT	CTA	CTT	AGC	AAG	GGT	403
	Val	Glu	Asn	Lys	Leu	Val	Glu	Leu	Glu	His	Thr	Leu	Leu	Ser	Lys	Gly	
				90					95					100			
30	TTC	AGA	GGT	GCA	TCA	CCT	CAC	ÇGG	AAA	TCC	ACC	TAA	AAGC	GTA (CAGG		450
	Phe	Arg	Gly	Ala	Ser	Pro	His	Arg	Lys	Ser	Thr						
			105					110									
	ATG'	TAAT	GCC .	AGTG	GTGG.	AA A'	CAT')AAA1	G A	CACT	FTGA	GTA	3				493
35																	
	(2)	INF	ORMA	TION	FOR	SEQ	ID 1	: 01	50:								
		(.	i) S	EQUE	NCE (CHAR	ACTE	RIST	ICS:								

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	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
5		
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
	(B) CELL KIND: Epidermoid carcinoma	
	(C) CELL LINE: KB	
10	(D) CLONE NAME: HP10428	
	(ix) SEQUENCE CHARACTERISTICS:	
	(A) CHARACTERIZATION CODE: CDS	
	(B) EXISTENCE POSITION: 288 1385	
15	(C) CHARACTERIZATION METHOD: E	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50:	
	AGATTCCGGC CTGGAGCTCC CAGGGCCGAG CAGACCTTGG GACCTGTGAG CGCTGCATCC	60
20	AATTAACCAT GGGAAGGGTC AGCACCAGCC ACCAGCCCCT TAGGTGAGGA CTCTGCCTGG	120
	GGCTCTGCTG ATGGTTCCGA ATCATGGAGC TGCAGAGAGC TCCTCCAGCC TGGAGACGTT	180
	CTTGGTGAAA GCTGTGGTCT AACTCCACCG GCTCTTCCTG CACATTGTAT TCAAGAGGGG	240
	TGCCTGCCCC CGCTGACTCA GGAGCTCCGG TGCTGCAGCC GCCACGA ATG GGG AGG	296
	Met Gly Arg	
25	1	344
	TGG GCC CTC GAT GTG GCC TTT TTG TGG AAG GCG GTG TTG ACC CTG GGG	244
	Trp Ala Leu Asp Val Ala Phe Leu Trp Lys Ala Val Leu Thr Leu Gly	
	5 10 15	392
20	CTG GTG CTT CTC TAC TAC TGC TTC TCC ATC GGC ATC ACC TTC TAC AAC	392
30	Leu Val Leu Leu Tyr Tyr Cys Phe Ser Ile Gly Ile Thr Phe Tyr Asn 20 25 30 35	
	20	440
	AAG TGG CTG ACA AAG AGC TTC CAT TTC CCC CTC TTC ATG ACG ATG CTG	440
	Lys Trp Leu Thr Lys Ser Phe His Phe Pro Leu Phe Met Thr Met Leu 40 45 50	
2 E	CAC CTG GCC GTG ATC TTC CTC TTC TCC GCC CTG TCC AGG GCG CTG GTT	488
35		700
	His Leu Ala Val Ile Phe Leu Phe Ser Ala Leu Ser Arg Ala Leu Val	

CAG TGC TCC AGC CAC AGG GCC CGT GTG GTG CTG AGC TGG GCC GAC TAC

	Gln	Cys		Ser	His	Arg	Ala		Val	Val	Leu	Ser	Trp	Ala	Asp	Tyr	
			70					75					80				
													CTT				584
	Leu	Arg	Arg	Val	Ala	Pro	Thr	Ala	Leu	Ala	Thr	Ala	Leu	Asp	Val	Gly	
5		85					90					95					
	TTG	TCC	AAC	TGG	AGC	TTC	CTG	TAT	GTC	ACC	GTC	TCG	CTG	TAC	ACA	ATG	632
	Leu	Ser	Asn	Trp	Ser	Phe	Leu	Tyr	Val	Thr	Val	Ser	Leu	Tyr	Thr	Met	
	100					105					110					115	
	ACC	AAA	TCC	TCA	GCT	GTC	CTC	TTC	ATC	TTG	ATC	TTC	TCT	CTG	ATC	TTC	680
10	Thr	Lys	Ser	Ser	Ala	Val	Leu	Phe	Ile	Leu	Ile	Phe	Ser	Leu	Ile	Phe	
					120					125					130		
	AAG	CTG	GAG	GAG	CTG	CGC	GCG	GCA	CTG	GTC	CTG	GTG	GTC	CTC	CTC	ATC	728
	Lys	Leu	Glu	Glu	Leu	Arg	Ala	Ala	Leu	Val	Leu	Va1	Val	Leu	Leu	Ile	
				135					140					145			
15	GCC	GGG	GGT	CTC	TTC	ATG	TTC	ACC	TAC	AAG	TCC	ACA	CAG	TTC	AAC	GTG	776
	Ala	Gly	Gly	Leu	Phe	Met	Phe	Thr	Tyr	Lys	Ser	Thr	Gln	Phe	Asn	Val	
			150					155					160				
	GAG	GGC	TTC	GCC	TTG	GTG	CTG	GGG	GCC	TCG	TTC	ATC	GGT	GGC	ATT	CGC	824
	Glu	Gly	Phe	Ala	Leu	Val	Leu	Gly	Ala	Ser	Phe	Ile	Gly	Gly	Ile	Arg	
20		165					170					175					
	TGG	ACC	CTC	ACC	CAG	ATG	CTC	CTG	CAG	AAG	GCT	GAA	CTC	GGC	CTC	CAG	872
	Trp	Thr	Leu	Thr	Gln	Met	Leu	Leu	Gln	Lys	Ala	Glu	Leu	Gly	Leu	Gln	
	180					185					190					195	
	AAT	ccc	ATC	GAC	ACC	ATG	TTC	CAC	CTG	CAG	CCA	CTC	ATG	TTC	CTG	GGG	920
25	Asn	Pro	Ile	Asp	Thr	Met	Phe	His	Leu	Gln	Pro	Leu	Met	Phe	Leu	Gly	
					200					205					210		
	CTC	TTC	CCT	CTC	TTT	GCT	GTA	TTT	GAA	GGT	CTC	CAT	TTG	TCC	ACA	TCT	968
	Leu	Phe	Pro	Leu	Phe	Ala	Val	Phe	Glu	Gly	Leu	His	Leu	Ser	Thr	Ser	
				215					220					225			
30	GAG	AAA	ATC	TTC	CGT	TTC	CAG	GAC	ACA	GGG	CTG	CTC	CTG	CGG	GTA	CTT	1016
	G1u	Lys	Ile	Phe	Arg	Phe	Gln	Asp	Thr	Gly	Leu	Leu	Leu	Arg	Val	Leu	
			230					235					240				
	GGG	AGC	CTC	TTC	CTT	GGC	GGG	ATT	CTC	GCC	TTT	GGT	TTG	GGC	TTC	TCT	1064
													Leu				
35	•	245					250					255					
	GAG		CTC	CTG	GTC	TCC		ACC	TCC	AGC	CTC		CTC	TCC	ATT	GCC	1112
													Leu				
	260		. =3			265	0			-	270	•				275	

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	GGC	ATT	TTT	AAG	GAA	GTC	TGC	ACT	TTG	CTG	TTG	GCA	GCT	CAT	CTG	CTG	1160
٠	Gly	Ile	Phe	Lys	Glu	Val	Cys	Thr	Leu	Leu	Leu	Ala	Ala	His	Leu	Leu	
					280					285					290		
	GGC	GAT	CAG	ATC	AGC	CTC	CTG	AAC	TGG	CTG	GGC	TTC	GCC	CTC	TGC	CTC	1208
5	Gly	Asp	Gln	Ile	Ser	Leu	Leu	Asn	Trp	Leu	Gly	Phe	Ala	Leu	Cys	Leu	
				295					300					305			
	TCG	GGA	ATA	TCC	CTC	CAC	GTT	GCC	CTC	AAA	GCC	CTG	CAT	TCC	AGA	GGT	1256
	Ser	Gly	Ile	Ser	Leu	His	Val	Ala	Leu	Lys	Ala	Leu	His	Ser	Arg	Gly	
			310					315					320				
10	GAT	GGT	GGC	ccc	AAG	GCC	TTG	AAG	GGG	CTG	GGC	TCC	AGC	CCC	GAC	CTG	1304
	Asp	Gly	Gly	Pro	Lys	Ala	Leu	Lys	Gly	Leu	Gly	Ser	Ser	Pro	Asp	Leu	
		325					330					335					
	GAG	CTG	CTG	CTC	CGG	AGC	AGC	CAG	CGG	GAG	GAA	GGT	GAC	AAT	GAG	GAG	1352
	Glu	Leu	Leu	Leu	Arg	Ser	Ser	Gln	Arg	Glu	Glu	Gly	Asp	Asn	Glu	Glu	
15	340					345					350					355	
	GAG	GAG	TAC	TTT	GTG	GCC	CAG	GGG	CAG	CAG	TGA	CCAG	CCA	GGGC.	TAAA		1400
	Glu	Glu	Tyr	Phe	Val	Ala	Gln	G1y	Gln	Gln							
					360					365							
																CGCCAG	1460
20																AGCCAG	1520
																CCAGGC	1580
																AGGGGA	1640
																AAGGCA	1700
																GGAGCT	1760
25																TAAAT	1820
																GACAGT	1880
																GTCTTA	1940
														TCC	CCAG	TGGGGC	2000
	CCC	ACTG	CAC	CTGC	TGGC	AG G	TAAA	AAAT	G AA	TGTT	TACT	GAG	T				2044
30																	

(2) INFORMATION FOR SEQ ID NO: 51:

35

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1043

(B) TYPE: Nucleic acid

(C) STRANDEDNESS: Double

(D) TOPOLOGY: Linear

(ii) SEQUENCE KIND: cDNA to mRNA

		(v	ri) C	RIGI	NAL	SOUF	RCE:										
				(A)	ORGA	NISN	1: H	omo s	sapie	ens							
				(B)	CELI	. KIN	ND: 5	Stoma	ch c	ance	er						
				(D)	CLO	E NA	ME:	HP10	1429								
5																	
		į)	(x) S	EQUE	ENCE	CHAI	RACTI	ERIST	rics:	:							
				(A)	CHAR	RACTI	ERIZA	OITA	1 COI	DE: C	DS						
				(B)	EXIS	STEN	CE PO	SITI	ON:	157.	. 83	37					
				(C)	CHAF	RACTI	ERIZA	OITA	N ME	CHOD:	E						
10															•		
		()	ci) S	EQUE	ENCE	DESC	CRIPT	: NOI	SEC) ID	NO:	51:					
																TTAGAC	60
																ATATTC	120
15	CTTT	TTTT	rgr (CTAC	AGA	AC T	TTTAT	TCCT	r GTC	SAAA							174
												Pro	Thr	Thr	•	Lys	
											1				5	4 m m	000
									TTC								222
	Thr	Leu	Met		Leu	Ser	Ser	Phe	Phe	Thr	Ser	Leu	GIY		rne	iie	
20	0.574		maa	10	4 mm	c m m	000	464	15	CC 4	mcc.	<u>ለ ጥር</u>	۸۵۵	20	۸۲۵	Δጥጥ	270
									CAA								270
	vai	116	25	261	TIE	Leu	Gly	30	Gln	MIA	ırþ	116	35	Der	1112	***	
	CCT	СТТ		GAC	ጥርጥ	CCT	TCA		GGG	AGC	АТТ	ттс		ACT	TAC	GGA	318
25									Gly								
		40	•••				45		,			50			•	·	
	CTT	TTT	CGT	GGG	GAG	AGT	AGT	GAA	GAA	TTG	AGT	CAC	GGA	CTT	GCA	GAA	366
									Glu								
	55			·		60					65					70	
30	CCA	AAG	AAA	AAG	TTT	GCA	GTT	TTA	GAG	ATA	CTG	AAT	AAT	TCT	TCC	CAA	414
	Pro	Lys	Lys	Lys	Phe	Ala	Val	Leu	Glu	Ile	Leu	Asn	Asn	Ser	Ser	Gln	
					75					80					85		
	AAA	ACT	CTG	CAT	TCG	GTG	ACT	ATC	CTG	TTC	CTG	GTC	CTG	AGT	TTG	ATC	462
	Lys	Thr	Leu	His	Ser	Val	Thr	Ile	Leu	Phe	Leu	Val	Leu	Ser	Leu	Ile	
35				90					95					100			
	ACG	TCG	CTG	CTG	AGC	TCT	GGG	TTT	ACC	TTC	TAC	AAC	AGC	ATC	AGC	AAC	510

Thr Ser Leu Leu Ser Ser Gly Phe Thr Phe Tyr Asn Ser Ile Ser Asn

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	CCT	TAC	CAG	ACA	TTC	CTG	GGG	CCG	ACG	GGG	GTG	TAC	ACC	TGG	AAC	GGG	558
	Pro	Tyr	Gln	Thr	Phe	Leu	Gly	Pro	Thr	Gly	Val	Tyr	Thr	Trp	Asn	Gly	
		120					125					130					
	CTC	GGT	GCA	TCC	TTC	GTT	TTT	GTG	ACC	ATG	ATA	CTG	TTT	GTG	GCG	AAC	606
5	Leu	Gly	Ala	Ser	Phe	Val	Phe	Val	Thr	Met	Ile	Leu	Phe	Val	Ala	Asn	
	135					140					145					150	
	ACG	CAG	TCC	AAC	CAA	CTC	TCC	GAA	GAG	TTG	TTC	CAA	ATG	CTT	TAC	CCG	654
	Thr	Gln	Ser	Asn	Gln	Leu	Ser	Glu	Glu	Leu	Phe	Gln	Met	Leu	Tyr	Pro	
					155					160					165		
10	GCA	ACC	ACC	AGT	AAA	GGA	ACG	ACC	CAC	AGT	TAC	GGA	TAC	TCG	TTC	TGG	702
	Ala	Thr	Thr	Ser	Lys	Gly	Thr	Thr	His	Ser	Tyr	Gly	Tyr	Ser	Phe	Trp	
				170					175					180			
	CTC	ATA	CTG	CTC	GTC	ATT	CTT	CTA	AAT	ATA	GTC	ACT	GTA	ACC	ATC	ATC	750
	Leu	Ile	Leu	Leu	Val	Ile	Leu	Leu	Asn	Ile	Val	Thr	Val	Thr	Ile	Ile	
15			185					190					195				
	ATT	TTC	TAC	CAG	AAG	GCC	AGA	TAC	CAG	CGG	AAG	CAG	GAG	CAG	AGA	AAG	798
	Ile	Phe	Tyr	Gln	Lys	Ala	Arg	Tyr	Gln	Arg	Lys	Gln	Glu	Gln	Arg	Lys	
		200					205					210					
	CCA	ATG	GAA	TAT	GCT	CCA	AGG	GAC	GGA	ATT	ATT	TTC	TGA	ATTC'	TCT '	TTCATC	850
20	Pro	Met	Glu	Tyr	Ala	Pro	Arg	Asp	Gly	Ile	Leu	Phe					
	215					220					225						
	TCA:	TTTT	GGC (STTG	CATC'	TA T	TGTA(CATC	A GC	CCTG	AGTA	GTA	ACTG	GTT A	AGCT'	TCTCTG	910
	GAC	AATT	CAG (CATG	GTAA	CG T	GACT	GTCA'	r cr	GTGA	CAGC	ATT'	TGTG'	TTT (CATG	ACACTG	970
	TGT	TCTT	CAT '	rgat(GCTG	TA C	TCCT	GAAA	A TT	TTTC	CCAC	AAG	GTTG	GG A	AAAT	GAATGG	1030
25	GAA	ATGT	CGC '	rgg													1043

(2) INFORMATION FOR SEQ ID NO: 52:

(i) SEQUENCE CHARACTERISTICS:

30 (A) LENGTH: 972

(B) TYPE: Nucleic acid

(C) STRANDEDNESS: Double

(D) TOPOLOGY: Linear

(ii) SEQUENCE KIND: cDNA to mRNA

35

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(B) CELL KIND: Liver

152

(D) CLONE NAME: HP10432

(ix) SEQUENCE CHARACTERISTICS:

5

(A) CHARACTERIZATION CODE: CDS

(B) EXISTENCE POSITION: 29.. 418

(C) CHARACTERIZATION METHOD: E

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:

10	AGACAGCGGC GGGCGCAGGA CGTGCACT ATG GCT CGG GGC TCG CTG CGC CGG	52
	Met Ala Arg Gly Ser Leu Arg Arg	
	1 5	
	TTG CTG CGG CTC CTC GTG CTG GGG CTC TGG CTG GCG TTG CTG C	100
	Leu Leu Arg Leu Leu Val Leu Gly Leu Trp Leu Ala Leu Leu Arg Ser	
15	10 15 20	
	GTG GCC GGG GAG CAA GCG CCA GGC ACC GCC CCC TGC TCC CGC GGC AGC	148
	Val Ala Gly Glu Gln Ala Pro Gly Thr Ala Pro Cys Ser Arg Gly Ser	
	25 30 35 40	
	TCC TGG AGC GCG GAC CTG GAC AAG TGC ATG GAC TGC GCG TCT TGC AGG	196
20	Ser Trp Ser Ala Asp Leu Asp Lys Cys Met Asp Cys Ala Ser Cys Arg	
	45 50 55	
	GCG CGA CCG CAC AGC GAC TTC TGC CTG GGC TGC GCT GCA GCA CCT CCT	244
	Ala Arg Pro His Ser Asp Phe Cys Leu Gly Cys Ala Ala Ala Pro Pro	
	60 65 70	
25	GCC CCC TTC CGG CTG CTT TGG CCC ATC CTT GGG GGC GCT CTG AGC CTG	292
	Ala Pro Phe Arg Leu Leu Trp Pro Ile Leu Gly Gly Ala Leu Ser Leu	
	75 80 85	
	ACC TTC GTG CTG GGG CTG CTT TCT GGC TTT TTG GTC TGG AGA CGA TGC	340
	Thr Phe Val Leu Gly Leu Leu Ser Gly Phe Leu Val Trp Arg Arg Cys	
30	90 95 100	
	CGC AGG AGA GAG AAG TTC ACC ACC CCC ATA GAG GAG ACC GGC GGA GAG	388
	Arg Arg Arg Glu Lys Phe Thr Thr Pro Ile Glu Glu Thr Gly Glu	
	105 110 115 120	
	GGC TGC CCA GCT GTG GCG CTG ATC CAG TGACA ATGT GCCCCCTGCC A CCGG	440
35	Gly Cys Pro Ala Val Ala Leu Ile Gln	
	125	
	GGCTCGCCCA CTCATCATTC ATTCATCCAT TCTAGAGCCA GTCTCTGCCT CCCAGACGCG	500
	GCGGGAGCCA AGCTCCTCCA ACCACAAGGG GGGTGGGGGG CGGTGAATCA CCTCTGAGGC	560

620

153

CTGGGCCCAG GGTTCAGGGG AACCTTCCAA GGTGTCTGGT TGCCCTGCCT CTGGCTCCAG

	AACAGAAAGG GAGCCTCACG CTGGCTCACA CAAAACAGCT GACACTGACT AAGGAACTGC	680
	AGCATTTGCA CAGGGGAGGG GGGTGCCCTC CTTCCTAGAG GCCCTGGGGG CCAGGCTGAC	740
	TTGGGGGGCA GACTTGACAC TAGGCCCCAC TCACTCAGAT GTCCTGAAAT TCCACCACGG	800
5	GGGTCACCCT GGGGGGTTAG GGACCTATTT TTAACACTAG GGGGCTGGCC CACTAGGAGG	860
	GCTGGCCCTA AGATACAGAC CCCCCAAACT CCCCAAAGCG GGGAGGAGAT ATTTATTTTG	920
	GGGAGAGTTT GGAGGGGAGG GAGAATTTAT TAATAAAAGA ATCTTTAACT TT	972
10	(2) INFORMATION FOR SEQ ID NO: 53:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 695	
	(B) TYPE: Nucleic acid	
	(C) STRANDEDNESS: Double	
15	(D) TOPOLOGY: Linear	
	(ii) SEQUENCE KIND: cDNA to mRNA	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Homo sapiens	
20	(B) CELL KIND: Liver	
	(C) CELL LINE:	
	(D) CLONE NAME: HP10433	
	(ix) SEQUENCE CHARACTERISTICS:	
25	(A) CHARACTERIZATION CODE: CDS	
	(B) EXISTENCE POSITION: 73 564	
	(C) CHARACTERIZATION METHOD: E	
	·	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:	
30	22242222422	60
	AAGATTTCAG CTGCGGGACG GTCAGGGGAG ACCTCCAGGC GCAGGGAAGG ACGGCCAGGG	
	TGACACGGAA GC ATG CGA CGG CTG CTG ATC CCT CTG GCC CTG TGG CTG GGC	111
	Met Arg Arg Leu Leu Ile Pro Leu Ala Leu Trp Leu Gly	
	1 5 10	159
35	GCG GTG GGC GTG GGC GTC GCC GAG CTC ACG GAA GCC CAG CGC CGG GGC	133
	Ala Val Gly Val Gly Val Ala Glu Leu Thr Glu Ala Gln Arg Gly	_
	15 20 25	207
	CTG CAG GTG GCC CTG GAG GAA TTT CAC AAG CAC CCG CCC GTG CAG TGG	207

	Leu	Gln	Val	Ala	Leu	Glu	G1u	Phe	His	Lys	His	Pro	Pro	Va1	G1n	Trp	
	30					35					40					45	
	GCC	TTC	CAG	GAG	ACC	AGT	GTG	GAG	AGC	GCC	GTG	GAC	ACG	CCC	TTC	CCA	255
	Ala	Phe	Gln	Glu	Thr	Ser	Val	Glu	Ser	Ala	Val	Asp	Thr	Pro	Phe	Pro	
5					50					55					60		
	GCT	GGA	ATA	TTT	GTG	AGG	CTG	GAA	TTT	AAG	CTG	CAG	CAG	ACA	AGC	TGC	303
	Ala	Gly	Ile	Phe	Val	Arg	Leu	Glu	Phe	Lys	Leu	Gln	Gln	Thr	Ser	Cys	
				65					70					75			
	CGG	AAG	AGG	GAC	TGG	AAG	AAA	CCC	GAG	TGC	AAA	GTC	AGG	CCC	AAT	GGG	351
10	Arg	Lys	Arg	Asp	Trp	Lys	Lys	Pro	G1u	Cys	Lys	Val	Arg	Pro	Asn	Gly	
			80					85					90				
	AGG	AAA	CGG	AAA	TGC	CTG	GCC	TGC	ATC	AAA	CTG	GGC	TCT	GAG	GAC	AAA	399
	Arg	Lys	Arg	Lys	Cys	Leu	Ala	Cys	Ile	Lys	Leu	Gly	Ser	Glu	Asp	Lys	
		95					100					105					
15	GTT	CTG	GGC	CGG	TTG	GTC	CAC	TGC	ccc	ATA	GAG	ACC	CAA	GTT	CTG	CGG	447
	Val	Leu	Gly	Arg	Leu	Val	His	Cys	Pro	Ile	Glu	Thr	Gln	Val	Leu	Arg	
	110					115					120					125	
									CAG								495
	Glu	.Ala	Glu	Glu	His	Gln	G1u	Thr	Gln	Cys	Leu	Arg	Val	Gln	Arg	Ala	
20					130					135					140		
									TTC								543
	Gly	Glu	Asp	Pro	His	Ser	Phe	Tyr	Phe	Pro	Gly	Gln	Phe	Ala	Phe	Ser	
				145					150					155			
	AAG	GCC	CTG	CCC	CGC	AGC	TAA	GCCA	GCA	CTGA	GCTG	CG T	GGTG(CCTC			590
25	Lys	Ala	Leu	Pro	Arg	Ser											
			160														650
														AGA (GGAC	CCCGTT	650
	CTA	TCCC	CAG	CCAT	GATA	AT A	AAGC	TGCT	C TC	CCAG	CTGC	CTC	TC				695
30																	
	(2)			TION													
		(i) S	EQUE					ics:								
				, ,		GTH:											
								ic a									
35				• •					oubl	е							
			• • -	, ,				inea		n							
		(11)	SEQU	ENCE	KIN	n: c	DNA	to m	KNA							

PCT/JP98/02445

(vi) ORIGINAL SOURCE:

				(A)	ORGA	NISM	: Но	mo s	apie	ns							
				(B)	CELL	KIN	D: S	toma	ch c	ance	r						
				(D)	CLON	E NA	ME:	HP10	480								
5																	
		(i	x) S	EQUE	NCE	CHAR	ACTE	RIST	ICS:								
				(A)	CHAR	ACTE	RIZA	TION	COD	E: C	DS						
				(B)	EXIS	TENC	E PO	SITI	ON:	80	661						
				(C)	CHAR	ACTE	RIZA	TION	MET	HOD:	E						
10																	
		(x	i) S	EQUE	NCE	DESC	RIPI	: NOI	SEQ	ID	NO:	54:					
																CTCGG	60
	cccc	GCGC	CG C	CCGT	'CAAC											TGC	112
15						Met	Ile	Arg	Cys	Gly	Leu	ı Ala	Cys	Glu		Cys	
						1				5				mm.c	10		160
													GCC				160
	Arg	Trp	Ile	Leu	Pro	Leu	Leu	Leu		Ser	Ala	Ile	Ala		Asp	TTE	
				15					20		mom	400	CAC	25	CCC	CAG	208
20													GAC				200
	Ile	Ala		Ala	Gly	Arg	Gly		Leu	Gin	ser	Ser	Asp 40	птэ	Gly	0111	
			30					35	mo o	C 4 4	CAC	ccc		GGC	AGC	GGG	256
													GGC Glv				
0.5	Thr		Ser	Leu	Trp	irp	ъуs 50	Cys	Ser	GIN	Olu	55	Gly	,			
25	mcc	45	CAC	CAG	GGC	ጥርጥ		AGC	СТС	ATG	GAG		GCG	TGG	GGT	AGA	304
													Ala				
	60	1)1	014	014	0.7	65					70	-				75	
		GCG	GCT	GCC	ATG	CTC	TTC	TGT	GGC	TTC	ATC	ATC	CTG	GTG	ATC	TGT	352
30													Leu				
					80			-		85					90		
	TTC	ATC	CTC	TCC	TTC	TTC	GCC	CTC	TGT	GGA	ccc	CAG	ATG	CTT	GTC	TTC	400
													Met				
				95					100					105			
35	CTG	AGA	GTG	ATT	GGA	GGT	CTC	CTT	GCC	TTG	GCT	GCT	GTG	TTC	CAG	ATC	448
																Ile	
			110					115					120				
	ATC	TCC	CTG	GTA	ATT	TAC	ccc	GTG	AAG	TAC	ACC	CAG	ACC	TTC	ACC	CTT	496

	Ile Ser I	Leu Val	Ile Ty	r Pro Va	l Lys T	Tyr Thr	Gln Thr	Phe Thr Leu	
	125			130			135		
	CAT GCC	AAC CGT	GCT GT	C ACT TA	C ATC T	TAT AAC	TGG GCC	TAC GGC TTT	544
	His Ala	Asn Arg	Ala Va	t Thr Ty	r Ile T	Tyr Asn	Trp Ala	Tyr Gly Phe	
5	140		14	5		150		155	
	GGG TGG	GCA GCC	ACG AT	T ATC CT	G ATC	GGC TGT	GCC TTC	TTC TTC TGC	592
	Gly Trp	Ala Ala	Thr Ile	e Ile Le	u Ile (Gly Cys	Ala Phe	Phe Phe Cys	
			160		1	165		170	
	TGC CTC	CCC AAC	TAC GA	A GAT GA	C CTT	CTG GGC	AAT GCC	AAG CCC AGG	640
10	Cys Leu	Pro Asn	Tyr Gl	u Asp As	p Leu I	Leu Gly	Asn Ala	Lys Pro Arg	
		175			180			185	
	TAC TTC	TAC ACA	TCT GC	C TA ACT	TGGG A	ATGAATG	rg ggagaa	AATC GCT	690
	Tyr Phe	Tyr Thr	Ser Al	В					
		190							
15	GCTGCTGA	GA TGGA	CTCCAG .	AAGAAGAA	AC TGT	TTCTCCA	GGCGACTI	TG AACCCATTTT	750
	TTGGCAGT	GT TCAT	ATTATT .	AAACTAGT	CA AAA	ATGCTAA	AATAATTI	GG GAGAAAATAT	810
	TTTTTAAG	TA GTGT	TATAGT	TTCATGTT	TA TCT	ATTATT	TGTTTTGT	GA AGTTGTGTCT	870
	TTTCACTA	ĄT TACC	TATACT .	ATGCCAAT	AT TTC	CTTATAT	CTATCCAT	AA CATTTATACT	930
	ACATTTGT.	AA GAGA	ATATGC .	ACGTGAAA	CT TAA	CACTTTA	TAAGGTAA	AAA ATGAGGTTTC	990
20	CAAGATTT	AA TAAT	CTGATC .	AAGTTCTT	GT TAT	TTCCAAA	TAGAATGG	AC TTGGTCTGTT	1050
	AAGGGCTA	AG GAGA	AGAGGA .	AGATAAGG	TT AAA	AGTTGTT	AATGACCA	AAA CATTCTAAAA	1110
	GAAATGCA	AAAA AA	AAGTTT	ATTTTCAA	GC CTT	CGAACTA	TTTAAGGA	AA GCAAAATCAT	1170
	TTCCTAAA	TG CATA	TCATTT	GTGAGAAT	TT CTC	ATAATA	TCCTGAAT	CA TTCATTTCAG	1230
	CTAAGGCT	TC ATGT	TGACTC	GATATGTC	AT CTA	GGAAAGT	ACTATTTC	CAT GGTCCAAACC	1290
25	TGTTGCCA	TA GTTG	GTAAGG	CTTTCCTI	TA AGT	GTGAAAT	ATTTAGAT	GA AATTTTCTCT	1350
	TTTAAAGT	TC TTTA	TAGGGT	TAGGG TG T	GG GAA	AATGCTA	TATTAATA	AAA TCTGTAGTGT	1410
	TTTGTGTT	TA TATG	TTCAGA	ACCAGAGT	AG ACT	GGATTGA	AAGATGGA	CT GGGTCTAATT	1470
	TATCATGA	CT GATA	GATCTG	GTTAAGTI	GT GTA	GTAAAGC	ATTAGGAG	GG TCATTCTTGT	153
	CACAAAAG	TG CCAC	TAAAAC	AGCCTCAG	GA GAA	TAAATGA	CTTGCTT	TTC TAAATCTCAG	159
30	GTTTATCT	GG GCTC	TATCAT	ATAGACAG	GC TTC	TGATAGT	TTGCAACT	GT AAGCAGAAAC	165
	CTACATAT	AG TTAA	AATCCT	GGTCTTTC	TT GGT.	AAACAGA	TTTTAAAT	GT CTGATATAAA	171
	ACATGCCA	CA GGAG	AATTCG	GGGATTTG	AG TTT	CTCTGAA	TAGCATA	TAT ATGATGCATC	177
	GGATAGGT	CA TTAT	GATTTT	TTACCATI	TC GAC	TTACATA	ATGAAAA	CCA ATTCATTTTA	183
	AATATCAG	TTAT TA	ATTTTG	TAAGTTGT	GG AAA	AAGCTAA	TTGTAGT	TTT CATTATGAAG	189
35	TTTTCCCA	AT AAAC	CAGGTA	TTCT					191

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CLAIMS

- A protein comprising an amino acid sequence selected from the group consisting of the amino acid sequences of SEQ
 ID NOS: 1 to 18.
 - 2. A DNA encoding the protein according to claim 1.
- 3. A cDNA comprising a nucleotide sequence selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 19 to 36.
 - 4. A cDNA according to claim 3, which comprises a nucleotide sequence selected from the group consisting of the nucleotide sequences of SEQ ID NOS: 37 to 54.
 - 5. An expression vector capable of in vitro translating the DNA according to any of claims 2 to 4 or expressing said DNA in an eukaryotic cell.

20

15

6. A transformed eukaryotic cell capable of expressing the DNA according to any of claims 2 to 4 to produce the protein according to claim 1.

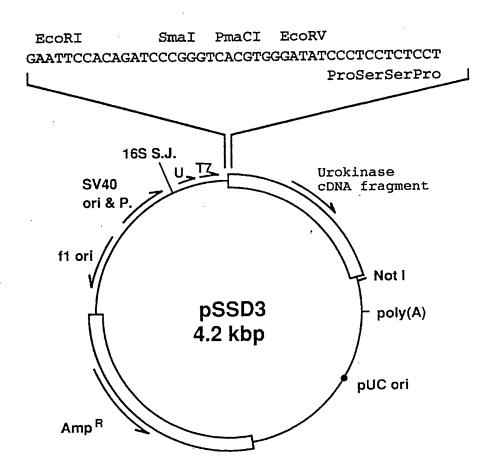


Fig.1

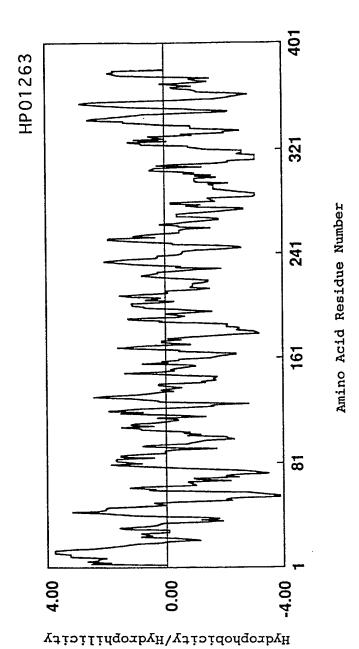
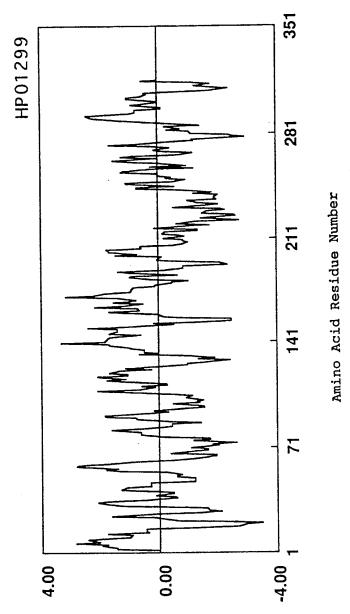


Fig.2



 ${\tt H} \lambda {\tt q} {\tt xobyoptc} {\tt r} {\tt f} \lambda {\tt t} {\tt d} {\tt q} {\tt xobyr} {\tt f} {\tt r} {\tt c} {\tt f} \lambda$

Fig.3

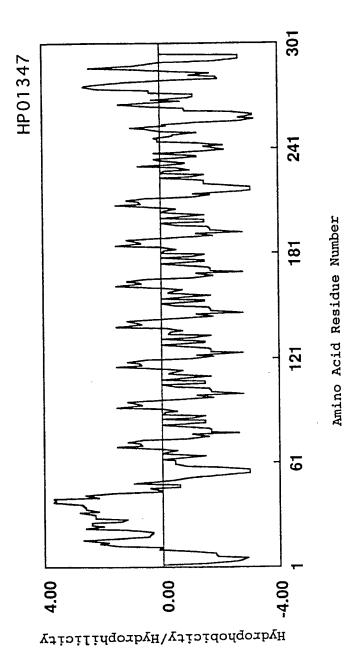


Fig.4

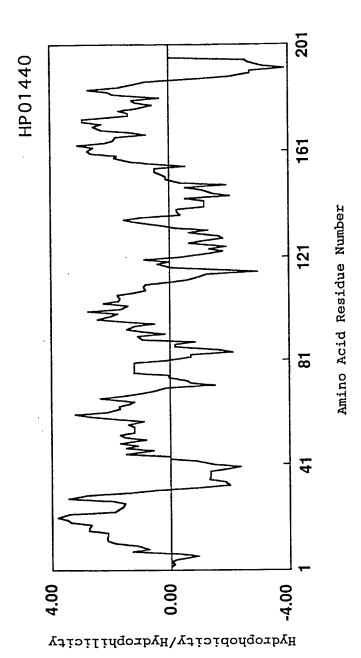
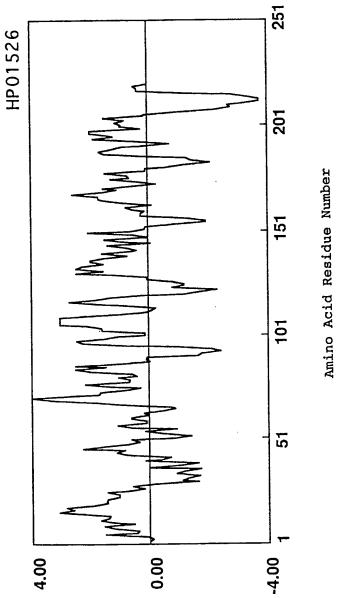


Fig.5



Ηλατοδρορτατελ\Ηλατοδριζιτετελ

Fig.6

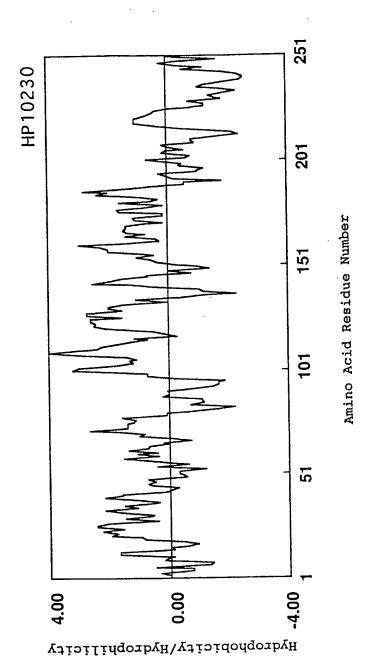


Fig.7

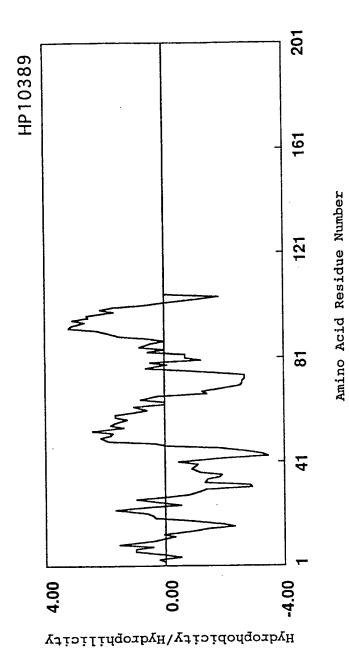


Fig.8

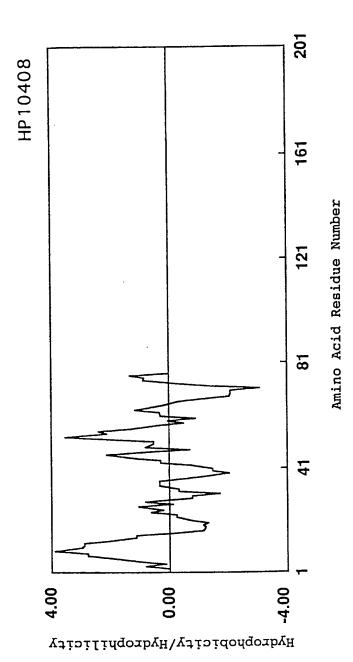


Fig.9

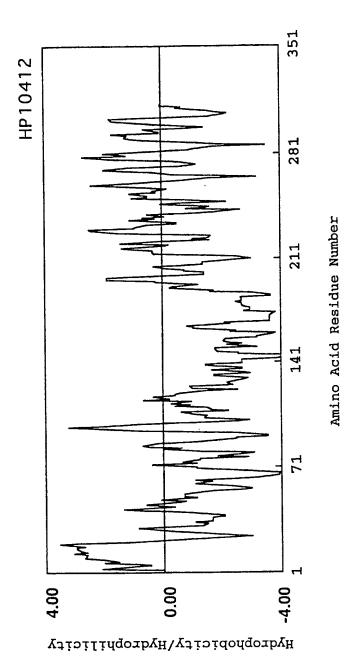


Fig.10

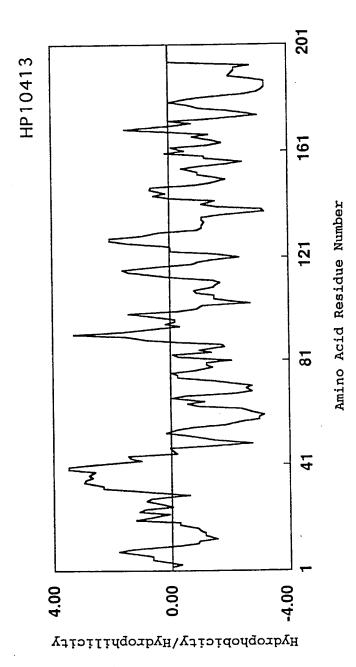
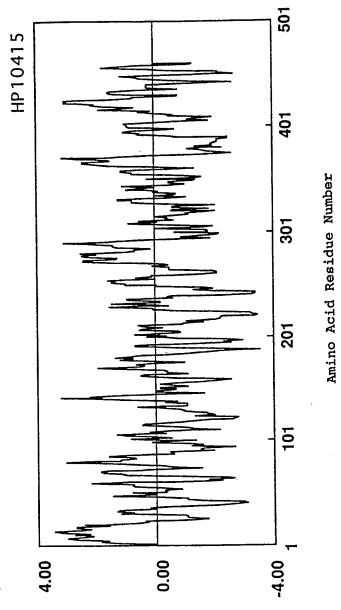


Fig.11



Ηλατορμορίς την Ηλατορμίζετ τη

Fig.12

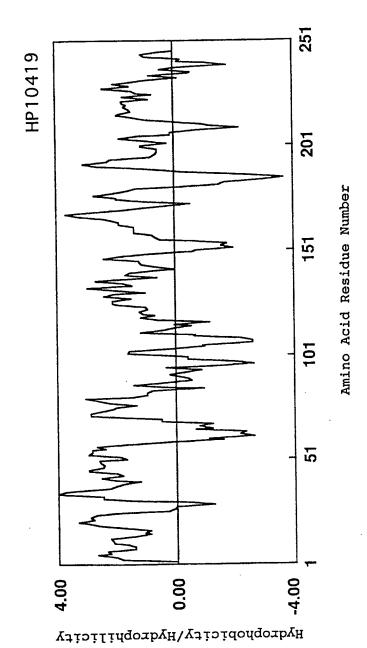


Fig.13

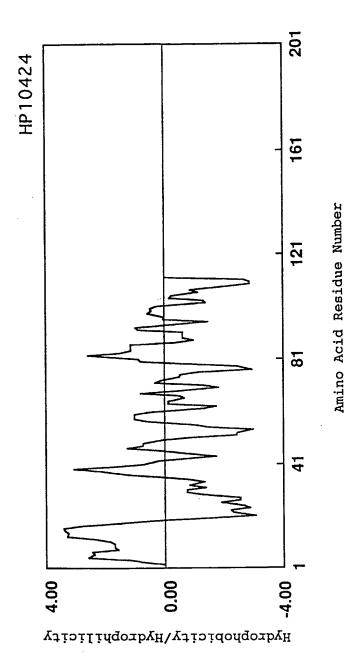
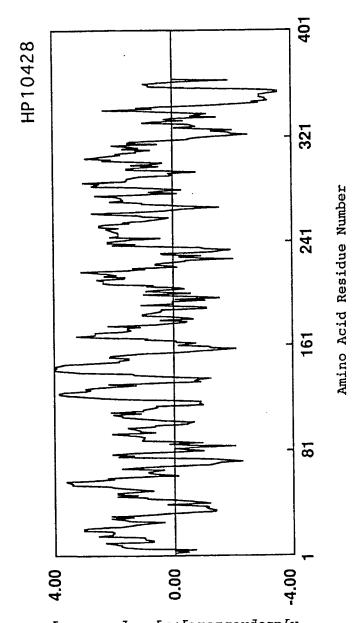


Fig.14



 ${\tt H}{\tt Aqrobyopicifh}{\tt H}{\tt Aqrobyijicifh}$

Fig.15

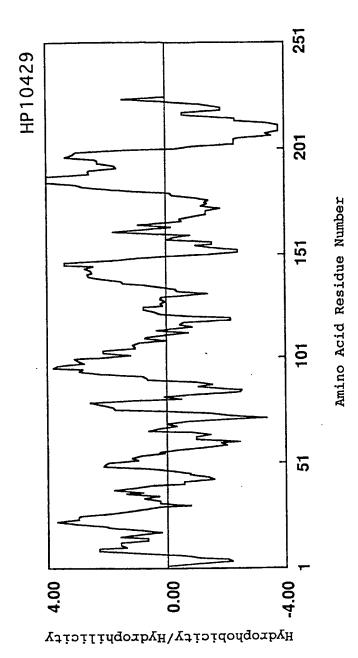


Fig.16

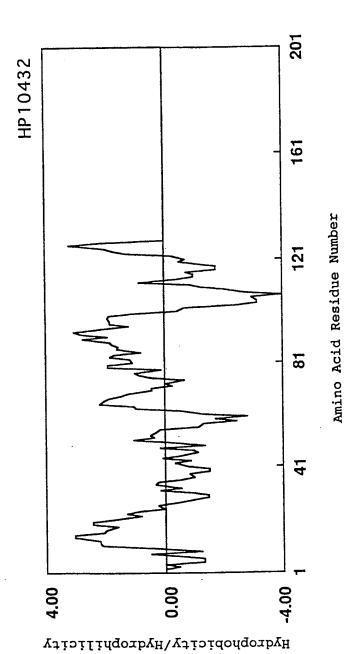


Fig.17

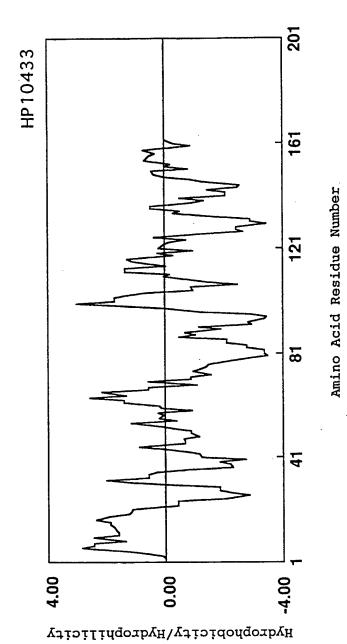


Fig.18

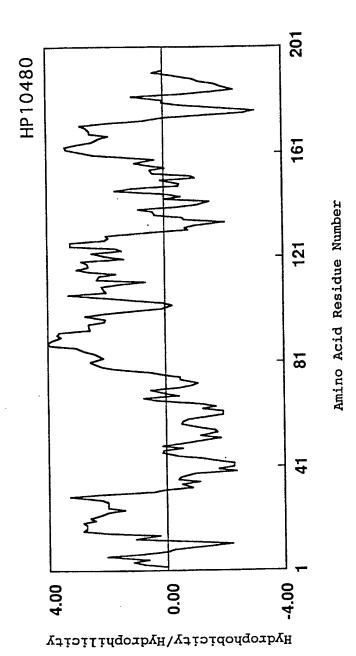


Fig.19

PCT

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51) International Patent Classification 6:		(11) International Publication Number: WO 98/5550
C12N 15/12, C07K 14/705, A61K 38/17, C12N 5/10, C12Q 1/37, C12N 9/72, 15/85	A3	(43) International Publication Date: 10 December 1998 (10.12.9)
21) International Application Number: PCT/JPS 22) International Filing Date: 3 June 1998 (C 30) Priority Data: 9/144948 3 June 1997 (03.06.97) 71) Applicants (for all designated States except US): S CHEMICAL RESEARCH CENTER [JP/JP] Nishi-Ohnuma 4-chome, Sagamihara-shi, K 229-0012 (JP). PROTEGENE INC. [JP/JP];	03.06.99 SAGAN ']; 4– Canagav	(AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, I LU, MC, NL, PT, SE). Published With international search report. Before the expiration of the time limit for amending the clar and to be republished in the event of the receipt of amendment of the international search report:
229-0014 (JP). SEKINE, Shingo [JP/JP]; II 101, 2-8-15, Atago, Ageo-shi, Saitama 362-00 YAMAGUCHI, Tomoko [JP/JP]; 5-13-11, T Katsushika-ku, Tokyo 125-0054 (JP). 74) Agents: AOYAMA, Tamotsu et al.; Aoyama & IMP Building, 3-7, Shiromi 1-chome, Chuo-ku, Oosaka 540-0001 (JP).	Kanagav Remon: 034 (JF Takasag Partne: Partne:	74 (14) (14) (14) (14) (14) (14) (14) (14
Proteins comprising any of the amino acid sequences of the nucelotide sequences of SEQ ID NOS: 19 to 36 are	of SEG provid	2 ID NOS: 1 to 18 and DNAs encoding said proteins and comprising a bd.

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INTERNATIONAL SEARCH REPORT

Internal Application No PCT/JP 98/02445

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A. CLASSIF IPC 6	C12N15/12 C07K14/705 A61K38/1 C12N9/72 C12N15/85	17 C12N5/10 C12Q1/37			
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	ion searched other than minimum documentation to the extent that s				
Electronic d	ata base consulted during the international search (name of data ba	se and, where practical, search terms used)			
C. DOCUME	ENTS CONSIDERED TO BE RELEVANT				
Category °	Citation of document, with indication, where appropriate, of the rel	evant passages Relevant to claim No.			
A	KYTE J. ET AL.: "A SIMPLE METHO DISPLAYING THE HYDROPATHIC CHARA PROTEIN" JOURNAL OF MOLECULAR BIOLOGY, vol. 157, no. 1, 5 May 1982, pag 105-132, XP000609692 cited in the application LIBERT F. ET AL.: "SELECTIVE AMPLIFICATION AND CLONING OF FOU MEMBERS OF THE G PROTEIN-COUPLED FAMILY" SCIENCE, vol. 244, 5 May 1989, pages 569-XP002041588	CTER OF A es R NEW RECEPTOR			
X Furti	her documents are listed in the continuation of box C.	Patent family members are listed in annex.			
° Special ca	tegories of cited documents :	Lander La			
"A" docume consic "E" earlier of filing of "L" docume which citatio "O" docume other: "P" docume later the consideration of the conside	ont defining the general state of the art which is not defining the general state of the art which is not dered to be of particular relevance document but published on or after the international	 *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention *X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone *Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. *&* document member of the same patent family Date of mailing of the international search report 			
Ì	2 September 1998	2 6. M. 99			
Name and r	mailing address of the ISA European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Macchia, G			

INTERNATIONAL SEARCH REPORT

Intern: hal Application No
PCT/JP 98/02445

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	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	MILLS A. AND DUGGAN M.J.: "ORPHAN SEVEN TRANSMEMBRANE DOMAIN RECEPTORS: REVERSING PHARMACOLOGY" TRENDS IN BIOTECHNOLOGY, vol. 12, February 1994, pages 47-49, XP002078287	
A	Database EMBL, entry Emest7:HS010272 Accession number N39010 25 January 1996 99% identity with Seq.ID:19 nt.647-1146. XP002078288 see the whole document	2-4
A	Database EMBL, entry Emest9:HS204207 Accession number H57204 7 October 1995 96% identity with Seq.ID:19 nt.1-437. XP002078292 cited in the application see the whole document	2-4

INTERNATIONAL SEARCH REPORT

In. ational application No. PCT/JP 98/02445

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-6: all partially (see subject 1, extra sheet)
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

1. Claims: 1-6 all partially.

A protein comprising an aminoacid sequence as in Seq.ID:1, encoding DNA, as in Seq.ID19 and 37, related expression vector and transformed eukaryotic cell.

2. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:2, 20 and 38.

3. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:3, 21 and 39.

4. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:4, 22 and 40.

5. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:5, 23 and 41.

6. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:6, 24 and 42.

7. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:7, 25 and 43.

8. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:8, 26 and 44.

9. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:9, 27 and 45.

10. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:10, 28 and 46.

11. Claims: 1-6 all partially.

As invention 1 but concerning Seq.ID:11, 29 and 47.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

- 12. Claims: 1-6 all partially.

 As invention 1 but concerning Seq.ID:12, 30 and 48.
- 13. Claims: 1-6 all partially.
 As invention 1 but concerning Seq.ID:13, 31 and 49.
- 14. Claims: 1-6 all partially.
 As invention 1 but concerning Seq.ID:14, 32 and 50.
- 15. Claims: 1-6 all partially.
 As invention 1 but concerning Seq.ID:15, 33 and 51.
- 16. Claims: 1-6 all partially.
 As invention 1 but concerning Seq.ID:16, 34 and 52.
- 17. Claims: 1-6 all partially.

 As invention 1 but concerning Seq.ID:17, 35 and 53.
- 18. Claims: 1-6 all partially.
 As invention 1 but concerning Seq.ID:18, 36 and 54.